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- (71) Applicant (*for all designated States except US*): **MERCK & CO. INC.** [US/US]; 126 East Lincoln Avenue, Rahway, NJ 07065-0907 (US).
- (72) Inventors; and
- (75) Inventors/Applicants (*for US only*): **COSFORD, Nicholas, D.** [GB/US]; 126 East Lincoln Avenue, Rahway, NJ 07065-0907 (US). **ROPPE, Jeffrey** [US/US]; 126 East Lincoln Avenue, Rahway, NJ 07065-0907 (US). **CHEN, Chixu** [US/US]; 126 East Lincoln Avenue, Rahway, NJ 07065-0907 (US). **SMITH, Nicholas** [US/US]; 126 East Lincoln Avenue, Rahway, NJ 07065-0907 (US). **REGER, Thomas** [US/US]; 126 East Lincoln Avenue, Rahway, NJ 07065-0907 (US).
- (74) Common Representative: **MERCK & CO. INC.**; 126 East Lincoln Avenue, Rahway, NJ 07065-0907 (US).
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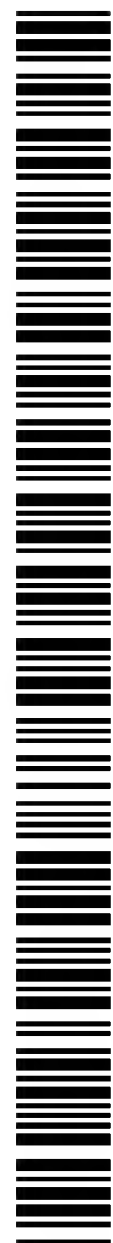
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(54) Title: HETEROARYL SUBSTITUTED TETRAZOLE MODULATORS OF METABOTROPIC GLUTAMATE RECEPTOR-5

(57) Abstract: Tetrazole compounds substituted directly, or by a bridge, with a heteroaryl moiety containing N adjacent to the point of connection of the heteroaryl, are mGluR5 modulators useful in the treatment of psychiatric and mood disorders such as, for example, schizophrenia, anxiety, depression, and panic, as well as in the treatment of pain and other diseases.



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TITLE OF THE INVENTION

HETEROARYL SUBSTITUTED TETRAZOLE MODULATORS OF
METABOTROPIC GLUTAMATE RECEPTOR-5

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BACKGROUND OF THE INVENTION

FIELD OF THE INVENTION

The present invention is directed to tetrazole compounds substituted
10 with a heteroaryl moiety. In particular, this invention is directed to tetrazole
compounds substituted directly, or by a bridge, with a heteroaryl moiety containing N
adjacent to the point of connection of the heteroaryl which are metabotropic glutamate
receptor – subtype 5 (“mGluR5”) modulators useful in the treatment of psychiatric
and mood disorders such as, for example, schizophrenia, anxiety, depression, and
15 panic, as well as in the treatment of pain, Parkinson’s disease, cognitive dysfunction,
epilepsy, drug addiction, drug abuse, drug withdrawal and other diseases

RELATED BACKGROUND

A major excitatory neurotransmitter in the mammalian nervous system
20 is the glutamate molecule, which binds to neurons, thereby activating cell surface
receptors. Such surface receptors are characterized as either ionotropic or
metabotropic glutamate receptors. The metabotropic glutamate receptors (“mGluR”)
are G protein-coupled receptors that activate intracellular second messenger systems
when bound to glutamate. Activation of mGluR results in a variety of cellular
25 responses. In particular, mGluR1 and mGluR5 activate phospholipase C, which is
followed by mobilizing intracellular calcium.

Modulation of metabotropic glutamate receptor subtype 5 (mGluR5) is
useful in the treatment of diseases that affect the nervous system (see for example
W.P.J.M Spooren et al., *Trends Pharmacol. Sci.*, 22:331-337 (2001) and references
30 cited therein). For example, recent evidence demonstrates the involvement of
mGluR5 in nociceptive processes and that modulation of mGluR5 using mGluR5-
selective compounds is useful in the treatment of various pain states, including acute,
persistent and chronic pain [K Walker et al., *Neuropharmacology*, 40:1-9 (2001); F.
Bordi, A. Ugolini *Brain Res.*, 871:223-233 (2001)], inflammatory pain [K Walker et

al., *Neuropharmacology*, 40:10-19 (2001); Bhavé et al. *Nature Neurosci.* 4:417-423 (2001)] and neuropathic pain [Dogrul et al. *Neurosci. Lett.* 292:115-118 (2000)].

Further evidence supports the use of modulators of mGluR5 in the treatment of psychiatric and neurological disorders. For example, mGluR5-selective compounds such as 2-methyl-6-(phenylethynyl)-pyridine ("MPEP") are effective in animal models of mood disorders, including anxiety and depression [W.P.J.M. Spooren et al., *J. Pharmacol. Exp. Ther.*, 295:1267-1275 (2000); E. Tatarczynska et al., *Brit. J. Pharmacol.*, 132:1423-1430 (2001); A. Klodzynska et al., *Pol. J. Pharmacol.*, 132:1423-1430 (2001)]. Gene expression data from humans indicate that modulation of mGluR5 may be useful for the treatment of schizophrenia [T. Ohnuma et al., *Mol. Brain. Res.*, 56:207-217 (1998); *ibid*, *Mol. Brain. Res.*, 85:24-31 (2000)]. Studies have also shown a role for mGluR5, and the potential utility of mGluR5-modulatory compounds, in the treatment of movement disorders such as Parkinson's disease [W.P.J.M. Spooren et al., *Europ. J. Pharmacol.* 406:403-410 (2000); H. Awad et al., *J. Neurosci.* 20:7871-7879 (2000); K. Ossawa et al. *Neuropharmacol.* 41:413-420 (2001)]. Other research supports a role for mGluR5 modulation in the treatment of cognitive dysfunction [G. Riedel et al., *Neuropharmacol.* 39:1943-1951 (2000)], epilepsy [A. Chapman et al., *Neuropharmacol.* 39:1567-1574 (2000)] and neuroprotection [V. Bruno et al., *Neuropharmacol.* 39:2223-2230 (2000)]. Studies with mGluR5 knockout mice and MPEP also suggest that modulation of these receptors may be useful in the treatment of drug addiction, drug abuse and drug withdrawal [C. Chiamulera et al. *Nature Neurosci.* 4:873-874 (2001)].

International Patent Publications WO 01/12627 and WO 99/26927 describe heteropolycyclic compounds and their use as metabotropic glutamate receptor antagonists.

U.S. Patent No. 3,647,809 describes pyridyl-1,2,4-oxadiazole derivatives. U.S. Patent No. 4,022,901 describes 3-pyridyl-5-isothiocyanophenyl oxadiazoles. International Patent Publication WO 98/17652 describes oxadiazoles, WO 97/03967 describes various substituted aromatic compounds, and WO 94/22846 describes various heterocyclic compounds.

Compounds that include ringed systems are described by various investigators as effective for a variety of therapies and utilities. For example, International Patent Publication No. WO 98/25883 describes ketobenzamides as calpain inhibitors, European Patent Publication No. EP 811610 and U.S. Patent Nos. 5,679,712, 5,693,672 and 5,747,541 describe substituted benzoylguanidine sodium

channel blockers, and U.S. Patent No. 5,736,297 describes ring systems useful as a photosensitive composition.

However, there remains a need for novel compounds and compositions that therapeutically inhibit mGluR5 with minimal side effects.

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SUMMARY OF THE INVENTION

The present invention is directed to novel tetrazole compounds substituted directly, or by a bridge, with a heteroaryl moiety containing N adjacent to the point of connection of the heteroaryl, are mGluR5 modulators useful in the treatment of psychiatric and mood disorders such as, for example, schizophrenia, anxiety, depression, and panic, as well as in the treatment of pain, Parkinson's disease, cognitive dysfunction, epilepsy, drug addiction, drug abuse, drug withdrawal and other diseases. This invention also provides a pharmaceutical composition which includes an effective amount of the novel tetrazole compounds substituted with a heteroaryl moiety, and a pharmaceutically acceptable carrier.

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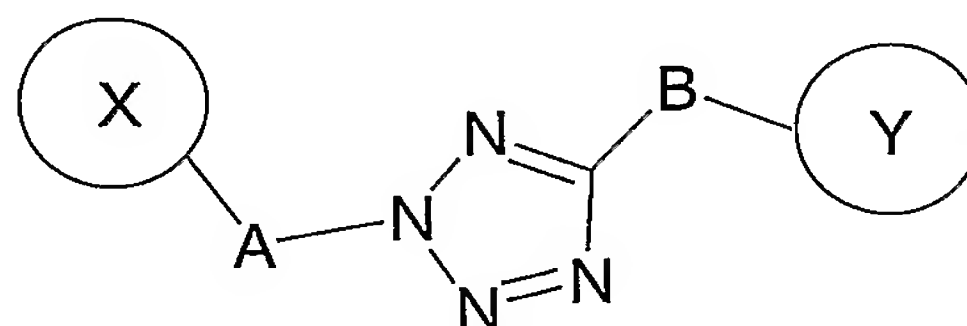
This invention further provides a method of treatment of psychiatric and mood disorders such as, for example, schizophrenia, anxiety, depression, and panic, as well as a method of treatment of pain, Parkinson's disease, cognitive dysfunction, epilepsy, drug addiction, drug abuse and drug withdrawal by the administration of an effective amount of the novel tetrazole compounds substituted with a heteroaryl moiety.

20

DETAILED DESCRIPTION OF THE INVENTION

The compounds of the present invention are represented by Formula

25 (I):



(I)

or a pharmaceutically acceptable salt thereof, wherein

X and Y each independently is aryl or heteroaryl wherein at least one of X and Y is a heteroaryl with N adjacent to the position of attachment to A or B respectively;

X is optionally substituted with 1-7 independent halogen, -CN, NO₂,
 5 -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -
 N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴,
 -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -
 C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form
 a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent,
 10 cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5
 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -
 O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or
 -N(C₀₋₆alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl,
 15 heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent
 halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -
 O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆
 alkyl)(aryl) substituents;

R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally
 20 substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇
 cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇
 cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂
 25 alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-
 NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

Y is optionally substituted with 1-7 independent halogen, -CN, NO₂,
 -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷,
 -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸,
 -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -
 30 C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form
 a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent,
 cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5
 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -
 O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or
 35 -N(C₀₋₆alkyl)(aryl) groups;

R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

In one aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is 2-pyridyl optionally substituted with 1-4 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -

O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

5 R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

10 Y is aryl or heteroaryl optionally substituted with 1-7 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are
15 combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

20 R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

25 R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

30 B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -

O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

5 In an embodiment of this one aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein
 X is 2-pyridyl optionally substituted with 1-4 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -
 C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -
 10 NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R²,
 -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are
 combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆
 6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further
 substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇
 15 7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇
 7cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent
 halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -
 20 O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆
 6alkyl)(aryl) substituents;

R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇
 7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇
 25 7cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

Y is phenyl optionally substituted with 1-5 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷,
 30 -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸,
 -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or
 -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to
 form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl

substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

5 R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

10 R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

15 B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

20 *R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

25 In a second aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

30 X is aryl or heteroaryl optionally substituted with 1-7 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

Y is 2-pyridyl optionally substituted with 1-4 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and
 any N may be an N-oxide.

In a third aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

Y is aryl or heteroaryl optionally substituted with 1-7 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶,

- NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵,
 -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein
 optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring
 fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl
 5 ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆
 alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆
 alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;
 R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl,
 heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent
 10 halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -
 O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆
 alkyl)(aryl) substituents;
 R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally
 substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇
 15 cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇
 cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;
 B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂
 alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-
 NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;
 20 R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl,
 heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent
 halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -
 O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆
 alkyl)(aryl) substituents; and
 25 any N may be an N-oxide.

In an embodiment of the third aspect of the invention, the compounds
 of this invention are represented by Formula (I) or a pharmaceutically acceptable salt
 30 thereof, wherein

X is phenyl optionally substituted with 1-5 independent halogen, -CN,
 NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -
 N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴,
 -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -

C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -
 5 O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -
 10 O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;
 15

A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

Y is 2-pyridyl optionally substituted with 1-4 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;
 20
 25

R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;
 30

R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

5 B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent
 10 halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

15

In a fourth aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is aryl or heteroaryl optionally substituted with 1-7 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -
 20 C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further
 25 substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent
 30 halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋

7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

Y is 1,3-thiazol-2-yl optionally substituted with 1-2 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

In an embodiment of this fourth aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent halogen, -CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁-6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀-6alkyl, -C₃-7cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), -N(C₀-6alkyl)(aryl) substituents;

R⁴ is -C₁-6alkyl, -C₃-7cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), -N(C₀-6alkyl)(aryl) substituents;

A is -C₀-4alkyl, -C₀-2alkyl-SO-C₀-2alkyl-, -C₀-2alkyl-SO₂-C₀-2alkyl-, -C₀-2alkyl-CO-C₀-2alkyl-, -C₀-2alkyl-NR⁹CO-C₀-2alkyl-, -C₀-2alkyl-NR¹SO₂-C₀-2alkyl- or -heteroC₀-4alkyl;

Y is 1,3-thiazol-2-yl optionally substituted with 1-2 independent halogen, -CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁-6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups;

R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

In a fifth aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is 3-pyridyl optionally substituted with 1-4 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -

O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

5 R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

10 Y is aryl or heteroaryl optionally substituted with 1-7 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are
15 combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

20 R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

25 R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

30 B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -

O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

5 In a sixth aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is aryl or heteroaryl optionally substituted with 1-7 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R²,
 10 -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

15 R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

20 R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

25 A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

30 Y is imidazolyl optionally substituted with 1-3 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋

6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

5 R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

10 R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

15 B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

20 R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

25 In an embodiment of this sixth aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

30 X is phenyl optionally substituted with 1-5 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) groups;

7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

Y is imidazolyl optionally substituted with 1-3 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

B is $-C_{0-4}alkyl$, $-C_{0-2}alkyl-SO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-SO_2-C_{0-2}alkyl-$, $-C_{0-2}alkyl-CO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-NR^{10}CO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-NR^1SO_2-C_{0-2}alkyl-$ or $-heteroC_{0-4}alkyl$;

R^9 and R^{10} each independently is $-C_{0-6}alkyl$, $-C_{3-7}cycloalkyl$,
 5 heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, $-CN$, $-C_{1-6}alkyl$, $-O(C_{0-6}alkyl)$, $-O(C_{3-7}cycloalkyl)$, $-O(aryl)$, $-O(heteroaryl)$, $-N(C_{0-6}alkyl)(C_{0-6}alkyl)$, $-N(C_{0-6}alkyl)(C_{3-7}cycloalkyl)$, $-N(C_{0-6}alkyl)(aryl)$ substituents; and

any N may be an N-oxide.

10

In a seventh aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is aryl or heteroaryl optionally substituted with 1-7 independent hydrogen, halogen, $-CN$, NO_2 , $-C_{1-6}alkyl$, $-C_{1-6}alkenyl$, $-C_{1-6}alkynyl$, $-OR^1$, $-NR^1R^2$, $-C(=NR^1)NR^2R^3$, $-N(=NR^1)NR^2R^3$, $-NR^1COR^2$, $-NR^1CO_2R^2$,
 15 $-NR^1SO_2R^4$, $-NR^1CONR^2R^3$, $-SR^4$, $-SOR^4$, $-SO_2R^4$, $-SO_2NR^1R^2$, $-COR^1$, $-CO_2R^1$, $-CONR^1R^2$, $-C(=NR^1)R^2$, or $-C(=NOR^1)R^2$ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the $-C_{1-6}alkyl$ substituent, cycloalkyl ring, or heterocycloalkyl
 20 ring each optionally is further substituted with 1-5 independent halogen, $-CN$, $-C_{1-6}alkyl$, $-O(C_{0-6}alkyl)$, $-O(C_{3-7}cycloalkyl)$, $-O(aryl)$, $-O(heteroaryl)$, $-N(C_{0-6}alkyl)(C_{0-6}alkyl)$, $-N(C_{0-6}alkyl)(C_{3-7}cycloalkyl)$, or $-N(C_{0-6}alkyl)(aryl)$ groups;

R^1 , R^2 , and R^3 each independently is $-C_{0-6}alkyl$, $-C_{3-7}cycloalkyl$, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent
 25 halogen, $-CN$, $-C_{1-6}alkyl$, $-O(C_{0-6}alkyl)$, $-O(C_{3-7}cycloalkyl)$, $-O(aryl)$, $-O(heteroaryl)$, $-N(C_{0-6}alkyl)(C_{0-6}alkyl)$, $-N(C_{0-6}alkyl)(C_{3-7}cycloalkyl)$, $-N(C_{0-6}alkyl)(aryl)$ substituents;

R^4 is $-C_{1-6}alkyl$, $-C_{3-7}cycloalkyl$, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, $-CN$, $-C_{1-6}alkyl$, $-O(C_{0-6}alkyl)$, $-O(C_{3-7}cycloalkyl)$, $-O(aryl)$, $-O(heteroaryl)$, $-N(C_{0-6}alkyl)(C_{0-6}alkyl)$, $-N(C_{0-6}alkyl)(C_{3-7}cycloalkyl)$, $-N(C_{0-6}alkyl)(aryl)$ substituents;
 30

A is $-C_{0-4}alkyl$, $-C_{0-2}alkyl-SO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-SO_2-C_{0-2}alkyl-$, $-C_{0-2}alkyl-CO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-NR^9CO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-NR^1SO_2-C_{0-2}alkyl-$ or $-heteroC_{0-4}alkyl$;

- Y is 1,3-oxazolyl optionally substituted with 1-2 independent hydrogen, halogen, -CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁-6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups; R⁵, R⁶, and R⁷ each independently is -C₀-6alkyl, -C₃-7cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), -N(C₀-6alkyl)(aryl) substituents;
- R⁸ is -C₁-6alkyl, -C₃-7cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), -N(C₀-6alkyl)(aryl) substituents;
- B is -C₀-4alkyl, -C₀-2alkyl-SO-C₀-2alkyl-, -C₀-2alkyl-SO₂-C₀-2alkyl-, -C₀-2alkyl-CO-C₀-2alkyl-, -C₀-2alkyl-NR¹⁰CO-C₀-2alkyl-, -C₀-2alkyl-NR¹SO₂-C₀-2alkyl- or -heteroC₀-4alkyl;
- R⁹ and R¹⁰ each independently is -C₀-6alkyl, -C₃-7cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), -N(C₀-6alkyl)(aryl) substituents; and
- any N may be an N-oxide.

- In an embodiment of this seventh aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

- X is phenyl optionally substituted with 1-5 independent hydrogen, halogen, -CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -

NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

Y is 1,3-oxazolyl optionally substituted with 1-2 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

5 B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent
10 halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

15 In an eighth aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is aryl or heteroaryl optionally substituted with 1-7 independent
hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R²,
20 -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent
30 halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

Y is isoxazolyl optionally substituted with 1-2 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

In an embodiment of this eighth aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

Y is isoxazolyl optionally substituted with 1-2 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

In a ninth aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is aryl or heteroaryl optionally substituted with 1-7 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -

O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

Y is pyrazolyl optionally substituted with 1-3 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -

O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

5 In an embodiment of this ninth aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

Y is pyrazolyl optionally substituted with 1-3 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋

6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

5 R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

10 R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

15 B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

25 In a tenth aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

30 X is aryl or heteroaryl optionally substituted with 1-7 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋

6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀-6alkyl, -C₃-7cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), -N(C₀-6alkyl)(aryl) substituents;

R⁴ is -C₁-6alkyl, -C₃-7cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), -N(C₀-6alkyl)(aryl) substituents;

A is -C₀-4alkyl, -C₀-2alkyl-SO-C₀-2alkyl-, -C₀-2alkyl-SO₂-C₀-2alkyl-, -C₀-2alkyl-CO-C₀-2alkyl-, -C₀-2alkyl-NR⁹CO-C₀-2alkyl-, -C₀-2alkyl-NR¹SO₂-C₀-2alkyl- or -heteroC₀-4alkyl;

Y is triazolyl optionally substituted with 1-3 independent hydrogen, halogen, -CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁-6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups;

R⁵, R⁶, and R⁷ each independently is -C₀-6alkyl, -C₃-7cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), -N(C₀-6alkyl)(aryl) substituents;

R⁸ is -C₁-6alkyl, -C₃-7cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), -N(C₀-6alkyl)(aryl) substituents;

B is $-C_{0-4}alkyl$, $-C_{0-2}alkyl-SO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-SO_2-C_{0-2}alkyl-$, $-C_{0-2}alkyl-CO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-NR^{10}CO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-NR^1SO_2-C_{0-2}alkyl-$ or $-heteroC_{0-4}alkyl$;

R^9 and R^{10} each independently is $-C_{0-6}alkyl$, $-C_{3-7}cycloalkyl$, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, $-CN$, $-C_{1-6}alkyl$, $-O(C_{0-6}alkyl)$, $-O(C_{3-7}cycloalkyl)$, $-O(aryl)$, $-O(heteroaryl)$, $-N(C_{0-6}alkyl)(C_{0-6}alkyl)$, $-N(C_{0-6}alkyl)(C_{3-7}cycloalkyl)$, $-N(C_{0-6}alkyl)(aryl)$ substituents; and
any N may be an N-oxide.

10

In an embodiment of this tenth aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent hydrogen, halogen, $-CN$, NO_2 , $-C_{1-6}alkyl$, $-C_{1-6}alkenyl$, $-C_{1-6}alkynyl$, $-OR^1$, $-NR^1R^2$, $-C(=NR^1)NR^2R^3$, $-N(=NR^1)NR^2R^3$, $-NR^1COR^2$, $-NR^1CO_2R^2$, $-NR^1SO_2R^4$, $-NR^1CONR^2R^3$, $-SR^4$, $-SOR^4$, $-SO_2R^4$, $-SO_2NR^1R^2$, $-COR^1$, $-CO_2R^1$, $-CONR^1R^2$, $-C(=NR^1)R^2$, or $-C(=NOR^1)R^2$ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the $-C_{1-6}alkyl$ substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, $-CN$, $-C_{1-6}alkyl$, $-O(C_{0-6}alkyl)$, $-O(C_{3-7}cycloalkyl)$, $-O(aryl)$, $-O(heteroaryl)$, $-N(C_{0-6}alkyl)(C_{0-6}alkyl)$, $-N(C_{0-6}alkyl)(C_{3-7}cycloalkyl)$, or $-N(C_{0-6}alkyl)(aryl)$ groups;

R^1 , R^2 , and R^3 each independently is $-C_{0-6}alkyl$, $-C_{3-7}cycloalkyl$, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, $-CN$, $-C_{1-6}alkyl$, $-O(C_{0-6}alkyl)$, $-O(C_{3-7}cycloalkyl)$, $-O(aryl)$, $-O(heteroaryl)$, $-N(C_{0-6}alkyl)(C_{0-6}alkyl)$, $-N(C_{0-6}alkyl)(C_{3-7}cycloalkyl)$, $-N(C_{0-6}alkyl)(aryl)$ substituents;

R^4 is $-C_{1-6}alkyl$, $-C_{3-7}cycloalkyl$, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, $-CN$, $-C_{1-6}alkyl$, $-O(C_{0-6}alkyl)$, $-O(C_{3-7}cycloalkyl)$, $-O(aryl)$, $-O(heteroaryl)$, $-N(C_{0-6}alkyl)(C_{0-6}alkyl)$, $-N(C_{0-6}alkyl)(C_{3-7}cycloalkyl)$, $-N(C_{0-6}alkyl)(aryl)$ substituents;

A is $-C_{0-4}alkyl$, $-C_{0-2}alkyl-SO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-SO_2-C_{0-2}alkyl-$, $-C_{0-2}alkyl-CO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-NR^9CO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-NR^1SO_2-C_{0-2}alkyl-$ or $-heteroC_{0-4}alkyl$;

Y is triazolyl optionally substituted with 1-3 independent hydrogen, halogen, -CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁-6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups;

R⁵, R⁶, and R⁷ each independently is -C₀-6alkyl, -C₃-7cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), -N(C₀-6alkyl)(aryl) substituents;

R⁸ is -C₁-6alkyl, -C₃-7cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), -N(C₀-6alkyl)(aryl) substituents;

B is -C₀-4alkyl, -C₀-2alkyl-SO-C₀-2alkyl-, -C₀-2alkyl-SO₂-C₀-2alkyl-, -C₀-2alkyl-CO-C₀-2alkyl-, -C₀-2alkyl-NR¹⁰CO-C₀-2alkyl-, -C₀-2alkyl-NR¹SO₂-C₀-2alkyl- or -heteroC₀-4alkyl;

R⁹ and R¹⁰ each independently is -C₀-6alkyl, -C₃-7cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), -N(C₀-6alkyl)(aryl) substituents; and

any N may be an N-oxide.

In an eleventh aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is aryl or heteroaryl optionally substituted with 1-7 independent hydrogen, halogen, -CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹,

- CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;
- 5 R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;
- 10 R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;
- 15 A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;
- Y is pyrimidinyl optionally substituted with 1-3 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;
- 20 R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;
- 30

R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

5 B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent
10 halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

15 In an embodiment of this eleventh aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -
20 C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further
25 substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent
30 halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋

7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

Y is pyrimidinyl optionally substituted with 1-3 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

In a twelvth aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is aryl or heteroaryl optionally substituted with 1-7 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups; R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

Y is pyrazinyl optionally substituted with 1-3 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

In an embodiment of this twelvth aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent

halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

5 R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

10 A is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR⁹CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

Y is pyrazinyl optionally substituted with 1-3 independent hydrogen, halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶,
15 -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;
20

R⁵, R⁶, and R⁷ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;
25

R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;
30

B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent

halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

5

Thus, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is 2-pyridyl optionally substituted with 1-4 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups.

Further, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is 2-pyridyl optionally substituted with 1-4 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups; and

Y is phenyl optionally substituted with 1-5 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -

C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -
 5 O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups.

Still further, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

10 X is aryl or heteroaryl optionally substituted with 1-7 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are
 15 combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups; and

20 Y is 2-pyridyl optionally substituted with 1-4 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are
 25 combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups.

30

Even further, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -

$N(=NR^1)NR^2R^3$, $-NR^1COR^2$, $-NR^1CO_2R^2$, $-NR^1SO_2R^4$, $-NR^1CONR^2R^3$, $-SR^4$,
 $-SOR^4$, $-SO_2R^4$, $-SO_2NR^1R^2$, $-COR^1$, $-CO_2R^1$, $-CONR^1R^2$, $-C(=NR^1)R^2$, or $-$
 $C(=NOR^1)R^2$ substituents, wherein optionally two substituents are combined to form
a cycloalkyl or heterocycloalkyl ring fused to X; wherein the $-C_{1-6}$ alkyl substituent,
5 cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5
independent halogen, $-CN$, $-C_{1-6}$ alkyl, $-O(C_{0-6}$ alkyl), $-O(C_{3-7}$ cycloalkyl), $-$
 $O(aryl)$, $-O(heteroaryl)$, $-N(C_{0-6}$ alkyl)(C_{0-6} alkyl), $-N(C_{0-6}$ alkyl)(C_{3-7} cycloalkyl), or
 $-N(C_{0-6}$ alkyl)(aryl) groups.

10 Still further, the compounds of this invention are represented by
Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent halogen, $-CN$,
 NO_2 , $-C_{1-6}$ alkyl, $-C_{1-6}$ alkenyl, $-C_{1-6}$ alkynyl, $-OR^1$, $-NR^1R^2$, $-C(=NR^1)NR^2R^3$, $-$
 $N(=NR^1)NR^2R^3$, $-NR^1COR^2$, $-NR^1CO_2R^2$, $-NR^1SO_2R^4$, $-NR^1CONR^2R^3$, $-SR^4$,
15 $-SOR^4$, $-SO_2R^4$, $-SO_2NR^1R^2$, $-COR^1$, $-CO_2R^1$, $-CONR^1R^2$, $-C(=NR^1)R^2$, or $-$
 $C(=NOR^1)R^2$ substituents, wherein optionally two substituents are combined to form
a cycloalkyl or heterocycloalkyl ring fused to X; wherein the $-C_{1-6}$ alkyl substituent,
cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5
independent halogen, $-CN$, $-C_{1-6}$ alkyl, $-O(C_{0-6}$ alkyl), $-O(C_{3-7}$ cycloalkyl), $-$
20 $O(aryl)$, $-O(heteroaryl)$, $-N(C_{0-6}$ alkyl)(C_{0-6} alkyl), $-N(C_{0-6}$ alkyl)(C_{3-7} cycloalkyl), or
 $-N(C_{0-6}$ alkyl)(aryl) groups; and

Y is 2-pyridyl optionally substituted with 1-4 independent halogen, $-$
 CN , NO_2 , $-C_{1-6}$ alkyl, $-C_{1-6}$ alkenyl, $-C_{1-6}$ alkynyl, $-OR^5$, $-NR^5R^6$,
 $-C(=NR^5)NR^6R^7$, $-N(=NR^5)NR^6R^7$, $-NR^5COR^6$, $-NR^5CO_2R^6$, $-NR^5SO_2R^8$, $-$
25 $NR^5CONR^6R^7$, $-SR^8$, $-SOR^8$, $-SO_2R^8$, $-SO_2NR^5R^6$, $-COR^5$, $-CO_2R^5$, $-CONR^5R^6$,
 $-C(=NR^5)R^6$, or $-C(=NOR^5)R^6$ substituents, wherein optionally two substituents are
combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the $-C_{1-6}$
alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further
substituted with 1-5 independent halogen, $-CN$, $-C_{1-6}$ alkyl, $-O(C_{0-6}$ alkyl), $-O(C_{3-7}$
30 $cycloalkyl)$, $-O(aryl)$, $-O(heteroaryl)$, $-N(C_{0-6}$ alkyl)(C_{0-6} alkyl), $-N(C_{0-6}$ alkyl)(C_{3-7}
 $cycloalkyl)$, or $-N(C_{0-6}$ alkyl)(aryl) groups.

And, the compounds of this invention are represented by Formula (I) or
a pharmaceutically acceptable salt thereof, wherein

Y is 1,3-thiazol-2-yl optionally substituted with 1-2 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups.

Further, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

Y is 1,3-thiazol-2-yl optionally substituted with 1-2 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups; and

X is phenyl optionally substituted with 1-5 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups.

As used herein, "alkyl" as well as other groups having the prefix "alk" such as, for example, alkoxy, alkanoyl, alkenyl, alkynyl and the like, means carbon chains which may be linear or branched or combinations thereof. Examples of alkyl groups include methyl, ethyl, propyl, isopropyl, butyl, sec- and tert-butyl, pentyl, 5 hexyl, heptyl and the like. "Alkenyl", "alkynyl" and other like terms include carbon chains containing at least one unsaturated C-C bond.

The term "cycloalkyl" means carbocycles containing no heteroatoms, and includes mono-, bi- and tricyclic saturated carbocycles, as well as fused ring systems. Such fused ring systems can include one ring that is partially or fully 10 unsaturated such as a benzene ring to form fused ring systems such as benzofused carbocycles. Cycloalkyl includes such fused ring systems as spirofused ring systems. Examples of cycloalkyl include cyclopropyl, cyclobutyl, cyclopentyl, cyclohexyl, decahydronaphthalene, adamantane, indanyl, indenyl, fluorenyl, 1,2,3,4-tetrahydronaphthalene and the like. Similarly, "cycloalkenyl" means carbocycles 15 containing no heteroatoms and at least one non-aromatic C-C double bond, and include mono-, bi- and tricyclic partially saturated carbocycles, as well as benzofused cycloalkenes. Examples of cycloalkenyl include cyclohexenyl, indenyl, and the like. §

The term "aryl" means an aromatic substituent which is a single ring or multiple rings fused together. When formed of multiple rings, at least one of the 20 constituent rings is aromatic. The preferred aryl substituents are phenyl and naphthyl groups.

The term "cycloalkyloxy" unless specifically stated otherwise includes a cycloalkyl group connected by a short C₁₋₂alkyl length to the oxy connecting atom.

The term "C₀₋₆alkyl" includes alkyls containing 6, 5, 4, 3, 2, 1, or no 25 carbon atoms. An alkyl with no carbon atoms is a hydrogen atom substituent when the alkyl is a terminal group and is a direct bond when the alkyl is a bridging group.

The term "hetero" unless specifically stated otherwise includes one or more O, S, or N atoms. For example, heterocycloalkyl and heteroaryl include ring systems that contain one or more O, S, or N atoms in the ring, including mixtures of 30 such atoms. The hetero atoms replace ring carbon atoms. Thus, for example, a heterocycloC₅alkyl is a five-member ring containing from 4 to no carbon atoms. Examples of heteroaryls include pyridinyl, quinolinyl, isoquinolinyl, pyridazinyl, pyrimidinyl, pyrazinyl, quinoxalinyl, furyl, benzofuryl, dibenzofuryl, thienyl, benzthienyl, pyrrolyl, indolyl, pyrazolyl, indazolyl, oxazolyl, benzoxazolyl, 35 isoxazolyl, thiazolyl, benzothiazolyl, isothiazolyl, imidazolyl, benzimidazolyl,

oxadiazolyl, thiadiazolyl, triazolyl, and tetrazolyl. Examples of heterocycloalkyls include azetidiny, pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, tetrahydrofuranyl, imidazolinyl, pyrrolidin-2-one, piperidin-2-one, and thiomorpholinyl.

5 The term "heteroC₀₋₄alkyl" means a heteroalkyl containing 3, 2, 1, or no carbon atoms. However, at least one heteroatom must be present. Thus, as an example, a heteroC₀₋₄alkyl having no carbon atoms but one N atom would be a -NH- if a bridging group and a -NH₂ if a terminal group. Analogous bridging or terminal groups are clear for an O or S heteroatom.

10 The term "amine" unless specifically stated otherwise includes primary, secondary and tertiary amines substituted with C₀₋₆alkyl.

 The term "carbonyl" unless specifically stated otherwise includes a C₀₋₆alkyl substituent group when the carbonyl is terminal.

15 The term "halogen" includes fluorine, chlorine, bromine and iodine atoms.

 The term "optionally substituted" is intended to include both substituted and unsubstituted. Thus, for example, optionally substituted aryl could represent a pentafluorophenyl or a phenyl ring. Further, optionally substituted multiple moieties such as, for example, alkylaryl are intended to mean that the aryl and the aryl groups are optionally substituted. If only one of the multiple moieties is optionally substituted then it will be specifically recited such as "an alkylaryl, the aryl optionally substituted with halogen or hydroxyl."

20 Compounds described herein contain one or more double bonds and may thus give rise to cis/trans isomers as well as other conformational isomers. The present invention includes all such possible isomers as well as mixtures of such isomers.

 Compounds described herein can contain one or more asymmetric centers and may thus give rise to diastereomers and optical isomers. The present invention includes all such possible diastereomers as well as their racemic mixtures, their substantially pure resolved enantiomers, all possible geometric isomers, and pharmaceutically acceptable salts thereof. The above Formula I is shown without a definitive stereochemistry at certain positions. The present invention includes all stereoisomers of Formula I and pharmaceutically acceptable salts thereof. Further, mixtures of stereoisomers as well as isolated specific stereoisomers are also included.

30 During the course of the synthetic procedures used to prepare such compounds, or in

using racemization or epimerization procedures known to those skilled in the art, the products of such procedures can be a mixture of stereoisomers.

The term "pharmaceutically acceptable salts" refers to salts prepared from pharmaceutically acceptable non-toxic bases or acids. When the compound of the present invention is acidic, its corresponding salt can be conveniently prepared from pharmaceutically acceptable non-toxic bases, including inorganic bases and organic bases. Salts derived from such inorganic bases include aluminum, ammonium, calcium, copper (ic and ous), ferric, ferrous, lithium, magnesium, manganese (ic and ous), potassium, sodium, zinc and the like salts. Particularly preferred are the ammonium, calcium, magnesium, potassium and sodium salts. Salts derived from pharmaceutically acceptable organic non-toxic bases include salts of primary, secondary, and tertiary amines, as well as cyclic amines and substituted amines such as naturally occurring and synthesized substituted amines. Other pharmaceutically acceptable organic non-toxic bases from which salts can be formed include ion exchange resins such as, for example, arginine, betaine, caffeine, choline, N,N'-dibenzylethylenediamine, diethylamine, 2-diethylaminoethanol, 2-dimethylaminoethanol, ethanolamine, ethylenediamine, N-ethylmorpholine, N-ethylpiperidine, glucamine, glucosamine, histidine, hydrabamine, isopropylamine, lysine, methylglucamine, morpholine, piperazine, piperidine, polyamine resins, procaine, purines, theobromine, triethylamine, trimethylamine, tripropylamine, tromethamine and the like.

When the compound of the present invention is basic, its corresponding salt can be conveniently prepared from pharmaceutically acceptable non-toxic acids, including inorganic and organic acids. Such acids include, for example, acetic, benzenesulfonic, benzoic, camphorsulfonic, citric, ethanesulfonic, fumaric, gluconic, glutamic, hydrobromic, hydrochloric, isethionic, lactic, maleic, malic, mandelic, methanesulfonic, mucic, nitric, pamoic, pantothenic, phosphoric, succinic, sulfuric, tartaric, p-toluenesulfonic acid and the like. Particularly preferred are citric, hydrobromic, hydrochloric, maleic, phosphoric, sulfuric, and tartaric acids.

The pharmaceutical compositions of the present invention comprise a compound represented by Formula I (or pharmaceutically acceptable salts thereof) as an active ingredient, a pharmaceutically acceptable carrier and optionally other therapeutic ingredients or adjuvants. Such additional therapeutic ingredients include, for example, i) opiate agonists or antagonists, ii) calcium channel antagonists, iii) 5HT receptor agonists or antagonists iv) sodium channel antagonists, v) NMDA

receptor agonists or antagonists, vi) COX-2 selective inhibitors, vii) NK1 antagonists, viii) non-steroidal anti-inflammatory drugs ("NSAID"), ix) GABA-A receptor modulators, x) dopamine agonists or antagonists, xi) selective serotonin reuptake inhibitors ("SSRI") and/or selective serotonin and norepinephrine reuptake inhibitors ("SSNRI"), xii) tricyclic antidepressant drugs, xiv) norepinephrine modulators, xv) L-DOPA, xvi) buspirone, xvii) lithium, xviii) valproate, ix) neurontin (gabapentin), xx) olanzapine, xxi) nicotinic agonists or antagonists including nicotine, xxii) muscarinic agonists or antagonists, xxiii) heroin substituting drugs such as methadone, levo-alpha-acetylmethadol, buprenorphine and naltrexone, and xxiv) disulfiram and acamprosate. The compositions include compositions suitable for oral, rectal, topical, and parenteral (including subcutaneous, intramuscular, and intravenous) administration, although the most suitable route in any given case will depend on the particular host, and nature and severity of the conditions for which the active ingredient is being administered. The pharmaceutical compositions may be conveniently presented in unit dosage form and prepared by any of the methods well known in the art of pharmacy.

* Creams, ointments, jellies, solutions, or suspensions containing the compound of Formula I can be employed for topical use. Mouth washes and gargles are included within the scope of topical use for the purposes of this invention.

Dosage levels from about 0.01mg/kg to about 140mg/kg of body weight per day are useful in the treatment of psychiatric and mood disorders such as, for example, schizophrenia, anxiety, depression, and panic, as well as being useful in the treatment of pain which are responsive to mGluR5 inhibition, or alternatively about 0.5mg to about 7g per patient per day. For example, schizophrenia, anxiety, depression, and panic may be effectively treated by the administration of from about 0.01mg to 75mg of the compound per kilogram of body weight per day, or alternatively about 0.5mg to about 3.5g per patient per day. Pain may be effectively treated by the administration of from about 0.01mg to 125mg of the compound per kilogram of body weight per day, or alternatively about 0.5mg to about 5.5g per patient per day. Further, it is understood that the mGluR5 inhibiting compounds of this invention can be administered at prophylactically effective dosage levels to prevent the above-recited conditions.

The amount of active ingredient that may be combined with the carrier materials to produce a single dosage form will vary depending upon the host treated and the particular mode of administration. For example, a formulation intended for

the oral administration to humans may conveniently contain from about 0.5mg to about 5g of active agent, compounded with an appropriate and convenient amount of carrier material which may vary from about 5 to about 95 percent of the total composition. Unit dosage forms will generally contain between from about 1mg to
5 about 1000mg of the active ingredient, typically 25mg, 50mg, 100mg, 200mg, 300mg, 400mg, 500mg, 600mg, 800mg or 1000mg.

It is understood, however, that the specific dose level for any particular patient will depend upon a variety of factors including the age, body weight, general health, sex, diet, time of administration, route of administration, rate of excretion,
10 drug combination and the severity of the particular disease undergoing therapy.

In practice, the compounds represented by Formula I, or pharmaceutically acceptable salts thereof, of this invention can be combined as the active ingredient in intimate admixture with a pharmaceutical carrier according to conventional pharmaceutical compounding techniques. The carrier may take a wide
15 variety of forms depending on the form of preparation desired for administration, e.g., oral or parenteral (including intravenous). Thus, the pharmaceutical compositions of the present invention can be presented as discrete units suitable for oral administration such as capsules, cachets or tablets each containing a predetermined amount of the active ingredient. Further, the compositions can be presented as a powder, as
20 granules, as a solution, as a suspension in an aqueous liquid, as a non-aqueous liquid, as an oil-in-water emulsion or as a water-in-oil liquid emulsion. In addition to the common dosage forms set out above, the compound represented by Formula I, or pharmaceutically acceptable salts thereof, may also be administered by controlled release means and/or delivery devices. The compositions may be prepared by any of
25 the methods of pharmacy. In general, such methods include a step of bringing into association the active ingredient with the carrier that constitutes one or more necessary ingredients. In general, the compositions are prepared by uniformly and intimately admixing the active ingredient with liquid carriers or finely divided solid carriers or both. The product can then be conveniently shaped into the desired
30 presentation.

Thus, the pharmaceutical compositions of this invention may include a pharmaceutically acceptable carrier and a compound or a pharmaceutically acceptable salt of Formula I. The compounds of Formula I, or pharmaceutically acceptable salts thereof, can also be included in pharmaceutical compositions in combination with one
35 or more other therapeutically active compounds.

The pharmaceutical carrier employed can be, for example, a solid, liquid, or gas. Examples of solid carriers include lactose, terra alba, sucrose, talc, gelatin, agar, pectin, acacia, magnesium stearate, and stearic acid. Examples of liquid carriers are sugar syrup, peanut oil, olive oil, and water. Examples of gaseous carriers include carbon dioxide and nitrogen.

In preparing the compositions for oral dosage form, any convenient pharmaceutical media may be employed. For example, water, glycols, oils, alcohols, flavoring agents, preservatives, coloring agents and the like may be used to form oral liquid preparations such as suspensions, elixirs and solutions; while carriers such as starches, sugars, microcrystalline cellulose, diluents, granulating agents, lubricants, binders, disintegrating agents, and the like may be used to form oral solid preparations such as powders, capsules and tablets. Because of their ease of administration, tablets and capsules are the preferred oral dosage units whereby solid pharmaceutical carriers are employed. Optionally, tablets may be coated by standard aqueous or nonaqueous techniques

A tablet containing the composition of this invention may be prepared by compression or molding, optionally with one or more accessory ingredients or adjuvants. Compressed tablets may be prepared by compressing, in a suitable machine, the active ingredient in a free-flowing form such as powder or granules, optionally mixed with a binder, lubricant, inert diluent, surface active or dispersing agent. Molded tablets may be made by molding in a suitable machine, a mixture of the powdered compound moistened with an inert liquid diluent. Each tablet preferably contains from about 0.1mg to about 500mg of the active ingredient and each cachet or capsule preferably containing from about 0.1mg to about 500mg of the active ingredient. Thus, a tablet, cachet, or capsule conveniently contains 0.1mg, 1mg, 5mg, 25mg, 50mg, 100mg, 200mg, 300mg, 400mg, or 500mg of the active ingredient taken one or two tablets, cachets, or capsules, once, twice, or three times daily.

Pharmaceutical compositions of the present invention suitable for parenteral administration may be prepared as solutions or suspensions of the active compounds in water. A suitable surfactant can be included such as, for example, hydroxypropylcellulose. Dispersions can also be prepared in glycerol, liquid polyethylene glycols, and mixtures thereof in oils. Further, a preservative can be included to prevent the detrimental growth of microorganisms.

Pharmaceutical compositions of the present invention suitable for injectable use include sterile aqueous solutions or dispersions. Furthermore, the compositions can be in the form of sterile powders for the extemporaneous preparation of such sterile injectable solutions or dispersions. In all cases, the final
5 injectable form must be sterile and must be effectively fluid for easy syringability. The pharmaceutical compositions must be stable under the conditions of manufacture and storage; thus, preferably should be preserved against the contaminating action of microorganisms such as bacteria and fungi. The carrier can be a solvent or dispersion medium containing, for example, water, ethanol, polyol (e.g. glycerol, propylene
10 glycol and liquid polyethylene glycol), vegetable oils, and suitable mixtures thereof.

Pharmaceutical compositions of the present invention can be in a form suitable for topical use such as, for example, an aerosol, cream, ointment, lotion, dusting powder, or the like. Further, the compositions can be in a form suitable for use in transdermal devices. These formulations may be prepared, utilizing a
15 compound represented by Formula I of this invention, or pharmaceutically acceptable salts thereof, via conventional processing methods. As an example, a cream or ointment is prepared by mixing hydrophilic material and water, together with about 5 wt% to about 10 wt% of the compound, to produce a cream or ointment having a desired consistency.

Pharmaceutical compositions of this invention can be in a form suitable for rectal administration wherein the carrier is a solid. It is preferable that the mixture forms unit dose suppositories. Suitable carriers include cocoa butter and other materials commonly used in the art. The suppositories may be conveniently formed by first admixing the composition with the softened or melted carrier(s)
25 followed by chilling and shaping in moulds.

In addition to the aforementioned carrier ingredients, the pharmaceutical formulations described above may include, as appropriate, one or more additional carrier ingredients such as diluents, buffers, flavoring agents, binders, surface-active agents, thickeners, lubricants, preservatives (including anti-oxidants)
30 and the like. Furthermore, other adjuvants can be included to render the formulation isotonic with the blood of the intended recipient. Compositions containing a compound described by Formula I, or pharmaceutically acceptable salts thereof, may also be prepared in powder or liquid concentrate form.

The compounds and pharmaceutical compositions of this invention
35 have been found to exhibit biological activity as mGluR5 inhibitors. Accordingly,

another aspect of the invention is the treatment in mammals of, for example, schizophrenia, anxiety, depression, and panic, pain, Parkinson's disease, cognitive dysfunction, epilepsy, drug addiction, drug abuse and drug withdrawal – maladies that are amenable to amelioration through inhibition of mGluR5 – by the
 5 administration of an effective amount of the compounds of this invention. The term “mammals” includes humans, as well as other animals such as, for example, dogs, cats, horses, pigs, and cattle. Accordingly, it is understood that the treatment of mammals other than humans is the treatment of clinical correlating afflictions to those above recited examples that are human afflictions.

10 Further, as described above, the compounds of this invention can be utilized in combination with other therapeutic compounds. In particular, the combinations of the mGluR5 inhibiting compound of this invention can be advantageously used in combination with i) opiate agonists or antagonists, ii) calcium channel antagonists, iii) 5HT receptor agonists or antagonists iv) sodium channel
 15 antagonists, v) NMDA receptor agonists or antagonists, vi) COX-2 selective inhibitors, vii) NK1 antagonists, viii) non-steroidal anti-inflammatory drugs (“NSAID”), ix) GABA-A receptor modulators, x) dopamine agonists or antagonists, xi) selective serotonin reuptake inhibitors (“SSRI”) and/or selective serotonin and norepinephrine reuptake inhibitors (“SSNRI”), xii) tricyclic antidepressant drugs, xiii)
 20 norepinephrine modulators, xiv) L-DOPA, xv) buspirone, xvi) lithium, xvii) valproate, xviii) neurontin (gabapentin), xix) olanzapine, xx) nicotinic agonists or antagonists including nicotine, xxi) muscarinic agonists or antagonists, xxii) heroin substituting drugs such as methadone, levo-alpha-acetylmethadol, buprenorphine and naltrexone, and xxiii) disulfiram and acamprosate.

25 The abbreviations used herein have the following tabulated meanings. Abbreviations not tabulated below have their meanings as commonly used unless specifically stated otherwise.

Ac	acetyl
AIBN	2,2'-azobis(isobutyronitrile)
BINAP	1,1'-bi-2-naphthol
Bn	benzyl
CAMP	cyclic adenosine-3',5'-monophosphate
DAST	(diethylamino)sulfur trifluoride

DEAD	diethyl azodicarboxylate
DBU	1,8-diazabicyclo[5.4.0]undec-7-ene
DIBAL	diisobutylaluminum hydride
DMAP	4-(dimethylamino)pyridine
DMF	N,N-dimethylformamide
Dppf	1,1'-bis(diphenylphosphino)-ferrocene
EDCI	1-(3-dimethylaminopropyl)-3-ethylcarbodiimide hydrochloride
Et ₃ N	triethylamine
GST	glutathione transferase
HMDS	hexamethyldisilazide
LDA	lithium diisopropylamide
m-CPBA	metachloroperbenzoic acid
MMPP	monoperoxyphthalic acid
MPPM	monoperoxyphthalic acid, magnesium salt 6H ₂ O
Ms	methanesulfonyl = mesyl = SO ₂ Me
MsO	methanesulfonate = mesylate
NBS	N-bromo succinimide
NSAID	non-steroidal anti-inflammatory drug
o-Tol	ortho-tolyl
OXONE®	2KHSO ₅ •KHSO ₄ •K ₂ SO ₄
PCC	pyridinium chlorochromate
Pd ₂ (dba) ₃	Bis(dibenzylideneacetone) palladium(0)
PDC	pyridinium dichromate
PDE	Phosphodiesterase
Ph	Phenyl
Phe	Benzenediyl
PMB	para-methoxybenzyl
Pye	Pyridinediyl
r.t.	room temperature
Rac.	Racemic
SAM	aminosulfonyl or sulfonamide or SO ₂ NH ₂
SEM	2-(trimethylsilyl)ethoxymethoxy

SPA	scintillation proximity assay
TBAF	tetra-n-butylammonium fluoride
Th	2- or 3-thienyl
TFA	trifluoroacetic acid
TFAA	trifluoroacetic acid anhydride
THF	Tetrahydrofuran
Thi	Thiophenediyl
TLC	thin layer chromatography
TMS-CN	trimethylsilyl cyanide
TMSI	trimethylsilyl iodide
Tz	1H (or 2H)-tetrazol-5-yl
XANTPHOS	4,5-Bis-diphenylphosphanyl-9,9-dimethyl-9H-xanthene
C ₃ H ₅	Allyl

ALKYL GROUP ABBREVIATIONS

Me	=	Methyl
Et	=	ethyl
<i>n</i> -Pr	=	normal propyl
<i>i</i> -Pr	=	isopropyl
<i>n</i> -Bu	=	normal butyl
<i>i</i> -Bu	=	isobutyl
<i>s</i> -Bu	=	secondary butyl
<i>t</i> -Bu	=	tertiary butyl
c-Pr	=	cyclopropyl
c-Bu	=	cyclobutyl
c-Pen	=	cyclopentyl
c-Hex	=	cyclohexyl

5

ASSAYS DEMONSTRATING BIOLOGICAL ACTIVITY

The compounds of this invention were tested against the hmGluR5a receptor stably expressed in mouse fibroblast Ltk⁻ cells (the hmGluR5a/L38-20 cell line) and activity was detected by changes in $[Ca^{++}]_i$, measured using the fluorescent Ca^{++} -sensitive dye, fura-2. InsP assays were performed in mouse fibroblast Ltk⁻ cells (LM5a cell line) stably expressing hmGluR5a. The assays described in International Patent Publication WO 0116121 can be used.

Calcium Flux Assay

The activity of compounds was examined against the hmGluR5a receptor stably expressed in mouse fibroblast Ltk⁻ cells (the hmGluR5a/L38 cell line). See generally Daggett et al., *Neuropharmacology* 34:871-886 (1995). Receptor activity was detected by changes in intracellular calcium ($[Ca^{2+}]_i$) measured using the fluorescent calcium-sensitive dye, fura-2. The hmGluR5a/L38-20 cells were plated onto 96-well plates, and loaded with 3 μ M fura-2 for 1h. Unincorporated dye was washed from the cells, and the cell plate was transferred to a 96-channel fluorimeter (SIBIA-SAIC, La Jolla, CA) which is integrated into a fully automated plate handling and liquid delivery system. Cells were excited at 350 and 385nm with a xenon source combined with optical filters. Emitted light was collected from the sample through a dichroic mirror and a 510nm interference filter and directed into a cooled CCD camera (Princeton Instruments). Image pairs were captured approximately every 1s, and ratio images were generated after background subtraction. After a basal reading of 20s, an EC₈₀ concentration of glutamate (10 μ M) was added to the well, and the response evaluated for another 60s. The glutamate-evoked increase in $[Ca]_i$ in the presence of the screening compound was compared to the response of glutamate alone (the positive control).

Phosphatidylinositol hydrolysis (PI) assays

Inositolphosphate assays were performed as described by Berridge et al. [Berridge et al, *Biochem. J.* 206: 587-5950 (1982); and Nakajima et al., *J. Biol. Chem.* 267:2437-2442 (1992)] with slight modifications. Mouse fibroblast Ltk cells expressing hmGluR5 (hmGluR5/L38- 20 cells) were seeded in 24-well plates at a density of 8x10⁵cells/well. One μ Ci of [³H]-inositol (Amersham PT6-271; Arlington Heights, Ill.; specific activity = 17.7 Ci/mmol) was added to each well and incubated for 16h at 37°C. Cells were washed twice and incubated for 45min in 0.5mL of standard Hepes buffered saline buffer (HBS; 125mM NaCl, 5mM KCl, 0.62mM

MgSO₄, 1.8mM CaCl₂, 20mM HEPES, 6mM glucose, pH to 7.4). The cells were washed with HBS containing 10mM LiCl, and 400μL buffer added to each well. Cells were incubated at 37°C for 20min. For testing, 50μL of 10X compounds used in the practice of the invention (made in HBS/LiCl (100mM)) was added and
5 incubated for 10 minutes. Cells were activated by the addition of 10μM glutamate, and the plates left for 1 hour at 37°C. The incubations were terminated by the addition of 1mL ice-cold methanol to each well. In order to isolate inositol phosphates (IPs), the cells were scraped from wells, and placed in numbered glass test tubes. One mL of chloroform was added to each tube, the tubes were mixed, and the
10 phases separated by centrifugation. IPs were separated on Dowex anion exchange columns (AG 1-X8 100-200 mesh formate form). The upper aqueous layer (750μL) was added to the Dowex columns, and the columns eluted with 3mL of distilled water. The eluents were discarded, and the columns were washed with 10mLs of 60mM ammonium formate/5mM Borax, which was also discarded as waste. Finally,
15 the columns were eluted with 4mL of 800mM ammonium formate/0.1M formic acid, and the samples collected in scintillation vials. Scintillant was added to each vial, and the vials shaken, and counted in a scintillation counter after 2 hours. Phosphatidylinositol hydrolysis in cells treated with certain exemplary compounds was compared to phosphatidylinositol hydrolysis in cells treated with the agonist
20 alone in the absence of compound.

The compounds of this application have mGluR5 inhibitory activity as shown by IC₅₀ values of less than 10 μM in the calcium flux assay or inhibition of
25 >50% at a concentration of 100 μM in the PI assay. Preferably, the compounds should have IC₅₀ values of less than 1 μM in the calcium flux assay and IC₅₀ values of less than 10 μM in the PI assay. Even more preferably, the compounds should have IC₅₀ values of less than 100 nM in the calcium flux assay and IC₅₀ values of less than 1 μM in the PI assay.

30 Examples 1-206 have mGluR5 inhibitory activity as shown by IC₅₀ values of 10 μM or better in the calcium flux assay and/or inhibition of >50% at 100 μM concentration in the PI assay

The examples that follow are intended as an illustration of certain preferred embodiments of the invention and no limitation of the invention is implied.

Unless specifically stated otherwise, the experimental procedures were performed under the following conditions. All operations were carried out at room or ambient temperature - that is, at a temperature in the range of 18-25°C. Evaporation of solvent was carried out using a rotary evaporator under reduced pressure (600-4000 pascals: 4.5-30 mm. Hg) with a bath temperature of up to 60°C. The course of reactions was followed by thin layer chromatography (TLC) and reaction times are given for illustration only. Melting points are uncorrected and 'd' indicates decomposition. The melting points given are those obtained for the materials prepared as described. Polymorphism may result in isolation of materials with different melting points in some preparations. The structure and purity of all final products were assured by at least one of the following techniques: TLC, mass spectrometry, nuclear magnetic resonance (NMR) spectrometry or microanalytical data. When given, yields are for illustration only. When given, NMR data is in the form of delta (δ) values for major diagnostic protons, given in parts per million (ppm) relative to tetramethylsilane (TMS) as internal standard, determined at 300 MHz, 400 MHz or 500 MHz using the indicated solvent. Conventional abbreviations used for signal shape are: s. singlet; d. doublet; t. triplet; m. multiplet; br. broad; etc. In addition, "Ar" signifies an aromatic signal. Chemical symbols have their usual meanings; the following abbreviations are used: v (volume), w (weight), b.p. (boiling point), m.p. (melting point), L (liter(s)), mL (milliliters), g (gram(s)), mg (milligrams(s)), mol (moles), mmol (millimoles), eq (equivalent(s)).

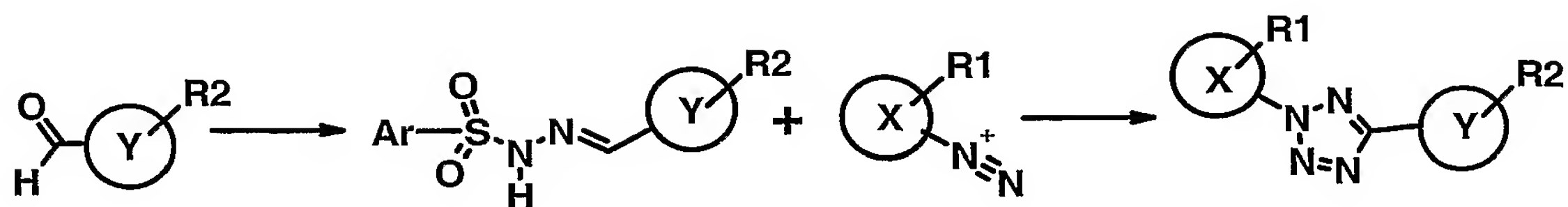
Methods of Synthesis

Compounds of the present invention can be prepared according to the following methods. The substituents are the same as in Formula I except where defined otherwise.

In accordance with another embodiment of the present invention, there are provided methods for the preparation of heteroaryl-substituted tetrazole compounds as described above. For example, many of the heterocyclic compounds described above can be prepared using synthetic chemistry techniques well known in the art (see *Comprehensive Heterocyclic Chemistry*, Katritzky, A. R. and Rees, C. W. eds., Pergamon Press, Oxford, 1984) from a heteroaryl-substituted tetrazole of Formula (I).

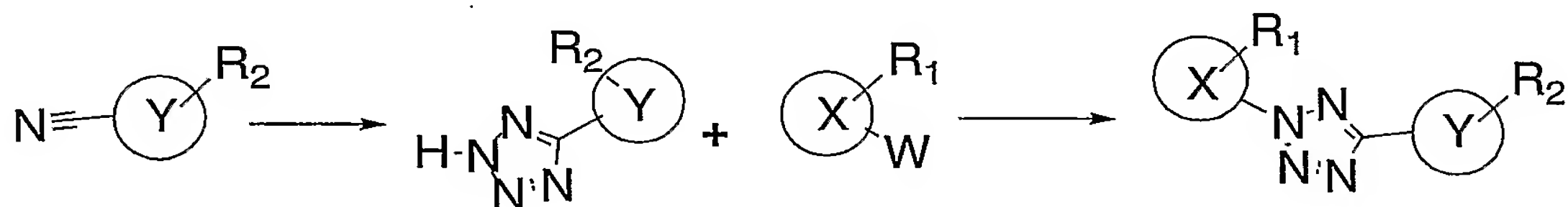
In Schemes 1 to 3 below, X, Y, R¹ and R² are as defined above for Formula (I).

Scheme 1



Referring to **Scheme 1**, ring system Y containing an aldehyde moiety (prepared using synthetic chemistry techniques well known in the art) is reacted with an arylsulfonylhydrazide in a suitable solvent (*e.g.* EtOH, MeOH, THF, DME, DMF *etc.*) at a temperature between 0°C to 100°C for 5 to 60min to form an arylsulfonylhydrazone. An amine-substituted X is treated with nitrous acid, at a temperature of -10°C to 0°C, in a suitable solvent such as, for example, water. In this manner an arenediazonium species is generated which then reacts with an arylsulfonylhydrazone in a 1,3-dipolar cycloaddition reaction to form a substituted tetrazole as shown (for example, see A.S. Shawali et al., *J. Heterocyclic Chem.* **1979**, 16, 123-128). The product from **Scheme 1**, a disubstituted tetrazole, can be isolated and purified employing standard techniques, such as solvent extraction, acid-base extraction, chromatography, crystallization, distillation and the like.

Scheme 2

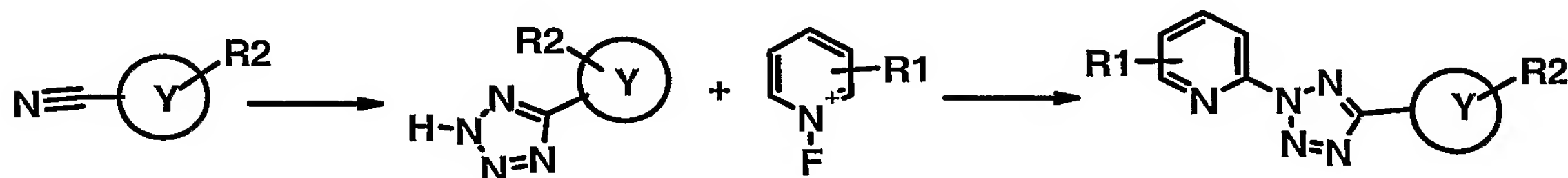


As shown in **Scheme 2** above, Y substituted with a nitrile functional group (prepared using methods well known in the art) is reacted with an azide moiety, such as LiN₃, NaN₃ or TMSN₃, in a suitable solvent (*e.g.* toluene, benzene, xylenes *etc.*) at a temperature in the range of about 25°C to 180°C to form a monosubstituted tetrazole. This reaction can conveniently be performed with an added catalyst such as dibutyltin oxide. The resulting tetrazole may then be coupled with X substituted with a group W. W maybe a metalloid species such as B(OR)₂, BiLn and the like and the reaction may be promoted with stoichiometric or catalytic amounts of metal salts such

as Cu(OAc)₂, CuI or CuOTf and the like. Conveniently, a base (*e.g.* pyridine, NEt₃, Cs₂CO₃, K₂CO₃ *etc.*) will also be present and the reaction is carried out in a suitable solvent (*e.g.* DCM, THF, DME toluene, MeCN, DMF, H₂O *etc.*). Additionally, molecular sieves may be used as a cocatalyst (see for example Fedorov, A. Y.; Finet, J-P. *Tetrahedron Lett.* **1999**, 40, 2747-2748).

Alternatively **W** may be a halogen or other functional group capable of undergoing a metal catalyzed *N*-arylation cross-coupling reaction in which case additional promoters such as 1,10-phenanthroline and dibenzylideneacetone may also be added to the reaction mixture. The cross-coupling reaction may be carried out at ambient temperature or heated to a temperature between about 30°C to 150°C. The reaction mixture is then maintained at a suitable temperature for a time in the range of about 4 to 72 hours, with 18 hours typically being sufficient (see for example Lam, P. Y. S.; Clark, C. G.; Saubern, S.; Adams, J.; Winters, M. P.; Cham, D. M. T.; Combs, A. *Tetrahedron Lett.* **1998**, 39, 2941-2944 and Kiyomori, A.; Marcoux, J. F.; Buchwald, S. L. *Tetrahedron Lett.* **1999**, 40, 2657-2660). The product from **Scheme 2**, a disubstituted tetrazole, can be isolated and purified employing standard techniques, such as solvent extraction, acid-base extraction, chromatography, crystallization, distillation and the like.

20 Scheme 3



Referring to **Scheme 3**, the monosubstituted tetrazole is prepared as described in **Scheme 2** from a suitable nitrile-substituted precursor. The tetrazole is then reacted with an *N*-fluoropyridinium salt, which may be optionally substituted, in the presence of a suitable base (*e.g.* MeONa, EtONa, *t*BuOK and the like) for a period of time sufficient for the reaction to proceed to completion, typically from about 1 to 12h, at a temperature in the range of about -100°C to 50°C, with -78°C to 23°C being advantageous (see for example Kiselyov, A. S. and Strekowski, L. *J. Heterocyclic Chem.* **1993**, 30, 1361-1364). The product from **Scheme 3**, a 2-pyridyltetrazole derivative, can be isolated and purified employing standard techniques, such as

solvent extraction, acid-base extraction, chromatography, crystallization, distillation and the like.

In addition, many of the heterocyclic intermediate compounds described above can be prepared using other synthetic chemistry techniques well known in the art (see *Comprehensive Heterocyclic Chemistry*, Katritzky, A. R. and Rees, C. W. eds., Pergamon Press, Oxford, 1984) and references cited there within.

EXAMPLE 1

2-[2-(3-chlorophenyl)-2H-tetrazol-5-yl]pyridine

3-Chloroaniline (762mg, 6.0mmol) was dissolved in 6N HCl (3.55mL, 21.3mmol), cooled to 0°C, and a solution of NaNO₂ (443mg, 6.42mmol) in H₂O (2mL) was added dropwise. The internal reaction temperature was maintained at <5°C by the addition of ice chips to the flask.

Separately, 2-pyridyl carboxaldehyde (643mg, 6.0mmol) and toluenesulfonyl hydrazide (1.12g, 6.0mmol) were combined in ethanol (15mL). The resulting reaction mixture was stirred at ambient temperature for 15min and monitored by TLC for the disappearance of aldehyde (2,4-DNP stain). NaOH pellets (1.2g, 30.0mmol) were then added along with H₂O (40mL) and the reaction mixture was cooled to 0°C with an ice bath.

The 3-chloroaniline diazotization reaction contents were then added dropwise *via* pipet. The resulting reaction was stirred for an additional 10min at 0°C, then the cooling bath was removed and the reaction was allowed to warm to ambient temperature. The crude mixture was diluted with EtOAc (300mL) and washed with H₂O (3x100mL). The organic phase was dried (MgSO₄), filtered, and concentrated *in vacuo*. The residue was chromatographed on silica gel eluting with hexanes:EtOAc (3:1) to afford 2-[2-(3-chlorophenyl)-2H-tetrazol-5-yl]pyridine as a pale orange solid.

¹H-NMR (CDCl₃, 300 MHz) δ 8.85 (d, J=4.9 Hz, 1H), 8.35 (m, 2H), 8.19 (d, J=7.29 Hz, 1H), 7.92 (dd, J=7.8 Hz, 7.7 Hz, 1H), 7.5 (m, 3H).

MS (ESI) 258.0 (M⁺+H).

EXAMPLE 2

3-(5-pyridin-2-yl-2H-tetrazol-2-yl)benzonitrile

By following the procedure described in **EXAMPLE 1** for the synthesis of 2-[2-(3-chlorophenyl)-2H-tetrazol-5-yl]pyridine, except that 3-amino benzonitrile (709mg, 6.0mmol) and 2-pyridyl carboxaldehyde (642mg, 6.0mmol)

were employed, 3-(5-pyridin-2-yl-2*H*-tetrazol-2-yl)benzonitrile was obtained as an orange solid.

¹H-NMR (CDCl₃, 300 MHz) δ 8.86 (d, J=3.9 Hz, 1H), 8.62 (s, 1H), 8.56 (d, J=8.2 Hz, 1H), 8.36 (d, J=7.9 Hz, 1H), 7.94 (dd, J=7.8 Hz, 7.7 Hz, 1H), 7.73-7.84 (m, 2H), 7.48 (m, 1H).

MS (ESI) 249.1 (M⁺+H).

EXAMPLE 3

2-(2-pyridin-3-yl-2*H*-tetrazol-5-yl)pyridine

Following the procedure described in **EXAMPLE 1** for the synthesis of 2-[2-(3-chlorophenyl)-2*H*-tetrazol-5-yl]pyridine, but using 3-amino pyridine (565mg, 6.0mmol) and 2-pyridyl carboxaldehyde (642mg, 6.0mmol), 2-(2-pyridin-3-yl-2*H*-tetrazol-5-yl)pyridine was obtained as an off white solid.

¹H-NMR (CDCl₃, 300 MHz) δ 9.57 (s, 1H), 8.86 (d, J=4.4 Hz, 1H), 8.79 (d, J=4.8 Hz, 1H), 8.57 (m, 1H), 8.36 (d, J=7.9 Hz, 1H), 7.93 (dd, J=7.8 Hz, 7.7 Hz, 1H), 7.58 (m, 1H), 7.48 (m, 1H).

MS (ESI) 225.1 (M⁺+H).

COMPOUND 1

(2-methyl-1,3-thiazol-4-yl)methanol

Ethyl 2-methyl-1,3-thiazole-4-carboxylate (15 g, 60mmol) was slurried in THF (40mL) and cooled to 0°C. Lithium aluminum hydride (60mL of a 1M solution in THF) was added slowly and the resulting reaction mixture was allowed to warm to 25°C. After 16h the reaction was quenched by the dropwise addition of water (2.28mL), 15% NaOH solution (2.28mL) and more water (6.84mL). Ethyl acetate (100mL) was added, the reaction mixture filtered, and the filtrate was concentrated *in vacuo*. The crude residue was chromatographed on silica gel with EtOAc:hexane (1:1) as eluant to afford (2-methyl-1,3-thiazol-4-yl)methanol as an oil.

¹H NMR (CD₃OD, 300 MHz) δ 7.21 (s, 1H), 4.53 (s, 2H), 2.67 (s, 3H).

COMPOUND 2

4-(2-methyl thiazole) carboxaldehyde

(2-Methyl-1,3-thiazol-4-yl)methanol (4.4g, 34mmol) was dissolved in CH₂Cl₂ (400mL). Magtrieve™ (44g) was added and the reaction was heated under reflux for 24h. The resulting mixture was filtered through Celite™, and the filter pad

was washed thoroughly with CH₂Cl₂. The filtrate was concentrated *in vacuo* to afford 2-methyl-1,3-thiazole-4-carbaldehyde as a yellow oil which was used in **EXAMPLE 4** without further purification.

¹H NMR (CDCl₃, 300 MHz) δ 9.98 (s, 1H), 8.06 (s, 1H), 2.79 (s, 3H).

5 MS (EI ionization) 127 (M⁺).

EXAMPLE 4

2-(3-chlorophenyl)-5-(2-methyl-1,3-thiazol-4-yl)-2H-tetrazole

Following the procedure described in **EXAMPLE 1** for the synthesis
10 of 2-[2-(3-chlorophenyl)-2H-tetrazol-5-yl]pyridine, 3-chloro aniline (565mg, 6.0mmol) and 4-(2-methyl thiazole) carboxaldehyde (508mg, 4.0mmol) were employed to obtain 2-(3-chlorophenyl)-5-(2-methyl-1,3-thiazol-4-yl)-2H-tetrazole as an orange solid.

¹H-NMR (CDCl₃, 300 MHz) δ 8.28 (s, 1H), 8.14 (m, 1H), 8.08 (s, 1H),
15 7.44-7.52 (m, 2H), 2.85 (s, 3H).

MS (ESI) 278.0 (M⁺+H).

EXAMPLE 5

3-[5-(2-methyl-1,3-thiazol-4-yl)-2H-tetrazol-2-yl]benzonitrile

Following the procedure described in **EXAMPLE 1** for the synthesis
20 of 2-[2-(3-chlorophenyl)-2H-tetrazol-5-yl]pyridine, 3-amino benzonitrile (651mg, 5.5mmol) and 4-(2-methyl thiazole) carboxaldehyde (700mg, 5.5mmol) were employed to obtain 3-[5-(2-methyl-1,3-thiazol-4-yl)-2H-tetrazol-2-yl]benzonitrile as an orange solid.

¹H-NMR (CDCl₃, 300 MHz) δ 8.60 (s, 1H), 8.53 (d, 1H), 8.13 (s, 1H),
25 7.68-7.82 (m, 2H).

MS (ESI) 269.0 (M⁺+H).

COMPOUND 3

5-(3-bromophenyl)-2H-tetrazole

3-Bromobenzonitrile (2.49g, 13.7mmol) and azidotrimethylsilane
(3.14g, 27.3mmol) were stirred in anhydrous toluene (100mL) under argon at ambient temperature. To this was added dibutyltin oxide (341mg, 1.37mmol) and the resulting reaction was fitted with a reflux condenser and heated with stirring at 110°C for 3h.
35 Upon cooling, the reaction mixture was concentrated *in vacuo*, and then concentrated

to dryness twice following the addition of MeOH (*ca.* 25mL each). The crude reaction mixture was dissolved in EtOAc (300mL) and washed with H₂O (2x100mL). The organic phase was washed with 1N aqueous NaOH (4x75mL), the combined basic aqueous portions were treated with 4N HCl to obtain an endpoint of pH=4 and the acidic aqueous phase was extracted with EtOAc (4x100mL washes). The combined EtOAc layers were dried (MgSO₄), filtered, and concentrated *in vacuo* to obtain 5-(3-bromophenyl)-2*H*-tetrazole as a white solid.

MS (ESI) 225.0 (M⁺+H).

10

EXAMPLE 6

2-[5-(3-bromophenyl)-2*H*-tetrazol-2-yl]pyridine

5-(3-Bromophenyl)-2*H*-tetrazole (448mg, 2.0mmol) was stirred in anhydrous MeOH (15mL) under argon at ambient temperature. To this was added NaOMe (4mL of 0.5M solution in MeOH, 2.0mmol), and the reaction flask was cooled to -78°C. A solution of *N*-fluoropyridinium triflate (331mg, 1.0mmol) in anhydrous MeOH (2mL) was then added dropwise *via* syringe. The resulting reaction mixture was stirred at -78°C for 30min, then warmed to ambient temperature and stirred for an additional 3h. The reaction mixture was partitioned between EtOAc (150mL) and 10% aqueous NaHCO₃ (50mL). The EtOAc layer was washed with additional 10% aqueous NaHCO₃ (2x50mL), and the combined aqueous layers were back extracted with EtOAc (100mL). The EtOAc layers were combined, dried (MgSO₄), filtered and concentrated *in vacuo*. The crude material was then chromatographed on silica gel eluting with hexanes:EtOAc (3:1) to afford 2-[5-(3-bromophenyl)-2*H*-tetrazol-2-yl]pyridine as a white solid.

¹H-NMR (CDCl₃, 300 MHz) δ 8.73 (d, 1H), 8.49 (s, 1H), 8.24 (m, 2H), 8.03 (dd, 1H), 7.64 (d, 1H), 7.52 (dd, 1H), 7.41 (dd, 1H).

MS (ESI) 324.0 (M⁺+Na).

30

EXAMPLE 7

2-[5-(3-chlorophenyl)-2*H*-tetrazol-2-yl]pyridine

Following the procedure in **COMPOUND 3** for the synthesis of intermediate 5-(3-bromophenyl)-2*H*-tetrazole, 3-chlorobenzonitrile (1.88g, 13.67mmol) was employed to obtain 5-(3-chlorophenyl)-2*H*-tetrazole.

MS (ESI) 181.1 (M⁺+H).

This material (500mg, 2.78mmol) was then carried on following the procedure in **EXAMPLE 6** for the synthesis of 2-[5-(3-bromophenyl)-2*H*-tetrazol-2-yl]pyridine to obtain 2-[5-(3-chlorophenyl)-2*H*-tetrazol-2-yl]pyridine as an off-white solid.

5 ¹H-NMR (CD₃OD, 300 MHz) δ 8.66 (d, 1H), 8.10-8.24 (m, 4H), 7.61 (dd, 1H), 7.40-7.52 (m, 2H).

MS (ESI) 280.0 (M⁺+Na).

EXAMPLE 8

10 **3-(2-pyridin-2-yl-2*H*-tetrazol-5-yl)benzonitrile**

Following the procedure in **COMPOUND 3** for the synthesis of 5-(3-bromophenyl)-2*H*-tetrazole, 1,3-dicyanobenzene (2.0 g, 15.63mmol) was employed to obtain 3-(2*H*-tetrazol-5-yl)benzonitrile.

MS (ESI) 172.1 (M⁺+H).

15 This material (921mg, 5.38mmol) was then carried on following the procedure in **EXAMPLE 6** for the synthesis of 2-[5-(3-bromophenyl)-2*H*-tetrazol-2-yl]pyridine to obtain 3-(2-pyridin-2-yl-2*H*-tetrazol-5-yl)benzonitrile as an off-white solid.

20 ¹H-NMR (CDCl₃, 300 MHz) δ 8.75 (d, 1H), 8.64 (s, 1H), 8.57 (d, 1H), 8.27 (d, 1H), 8.07 (dd, 1H), 7.81 (d, 1H), 7.68 (dd, 1H), 7.56 (dd, 1H).

MS (ESI) 271.0 (M⁺+Na).

EXAMPLE 9

2-[2-(3,5-difluorophenyl)-2*H*-tetrazol-5-yl]pyridine

25 Following the procedure described in **EXAMPLE 1** for the synthesis of 2-[2-(3-chlorophenyl)-2*H*-tetrazol-5-yl]pyridine, 3,5-difluoroaniline (131mg, 1.0mmol) and pyridine-2-carboxaldehyde (109mg, 1.0mmol) were employed to obtain 2-[2-(3,5-difluorophenyl)-2*H*-tetrazol-5-yl]pyridine as an orange solid.

30 ¹H-NMR (CDCl₃, 300 MHz) δ 8.87 (d, 1H), 8.37 (d, 1H), 7.90-7.97 (m, 3H), 7.46-7.51 (m, 1H), 6.97-7.03 (m, 1H).

MS (ESI) 260.0 (M⁺+H).

EXAMPLE 10

2-[2-(3-methoxyphenyl)-2*H*-tetrazol-5-yl]pyridine

Following the procedure described in **EXAMPLE 1** for the synthesis of 2-[2-(3-chlorophenyl)-2*H*-tetrazol-5-yl]pyridine, *m*-anisidine (135mg, 1.1mmol) and pyridine-2-carboxaldehyde (118mg, 1.1mmol) were employed to obtain 2-[2-(3-methoxyphenyl)-2*H*-tetrazol-5-yl]pyridine as an orange solid.

¹H-NMR (CDCl₃, 300 MHz) δ 8.84-8.86 (m, 1H), 8.34-8.38 (m, 1H), 7.80-7.94 (m, 3H), 7.43-7.50 (m, 2H), 7.04-7.07 (m, 1H), 3.93 (s, 3H).

MS (ESI) 254.0 (M⁺+H).

EXAMPLE 11

2-[2-(3-trifluoromethylphenyl)-2*H*-tetrazol-5-yl]pyridine

Following the procedure described in **EXAMPLE 1** for the synthesis of 2-[2-(3-chlorophenyl)-2*H*-tetrazol-5-yl]pyridine, 3-trifluoromethylaniline (195mg, 1.2mmol) and pyridine-2-carboxaldehyde (128 mg, 1.2mmol) were employed to obtain 2-[2-(3-trifluoromethylphenyl)-2*H*-tetrazol-5-yl]pyridine as an orange solid.

¹H-NMR (CDCl₃, 300 MHz) δ 8.87 (d, 1H), 8.60 (s, 1H), 8.52 (d, 1H), 8.39 (d, 1H), 7.91-7.97 (m, 1H), 7.72-7.81 (m, 2H), 7.46-7.50 (m, 1H).

MS (ESI) 292.0 (M⁺+H).

EXAMPLE 12

2-[2-(3-iodophenyl)-2*H*-tetrazol-5-yl]pyridine

Following the procedure described in **EXAMPLE 1** for the synthesis of 2-[2-(3-chlorophenyl)-2*H*-tetrazol-5-yl]pyridine, 3-iodoaniline (263mg, 1.2mmol) and pyridine-2-carboxaldehyde (128 mg, 1.2mmol) were employed to obtain 2-[2-(3-iodophenyl)-2*H*-tetrazol-5-yl]pyridine as an orange solid.

¹H-NMR (CDCl₃, 300 MHz) δ 8.85-8.87 (m, 1H), 8.68-8.69 (m, 1H), 8.26-8.38 (m, 2H), 7.84-7.95 (m, 2H), 7.44-7.49 (m, 1H), 7.27-7.35 (m, 1H).

MS (ESI) 350.0 (M⁺+H).

EXAMPLE 13

2-[2-(3-bromophenyl)-2*H*-tetrazol-5-yl]pyridine

Following the procedure described in **EXAMPLE 1** for the synthesis of 2-[2-(3-chlorophenyl)-2*H*-tetrazol-5-yl]pyridine, 3-bromoaniline (223mg, 1.3mmol) and pyridine-2-carboxaldehyde (135mg, 1.3mmol) were employed to obtain 2-[2-(3-bromophenyl)-2*H*-tetrazol-5-yl]pyridine as an orange solid.

$^1\text{H-NMR}$ (CDCl_3 , 300 MHz) δ 8.85-8.87 (m, 1H), 8.50-8.51 (m, 1H), 8.35-8.38 (m, 1H), 8.23-8.27 (m, 1H), 7.90-7.94 (m, 1H), 7.64-7.67 (m, 1H), 7.44-7.49 (m, 2H).

MS (ESI) 303.0 ($\text{M}^+ + \text{H}$).

5

EXAMPLE 14

2-[2-(3-Methylmercaptophenyl)-2*H*-tetrazol-5-yl]pyridine

Following the procedure described in **EXAMPLE 1** for the synthesis of 2-[2-(3-chlorophenyl)-2*H*-tetrazol-5-yl]pyridine, 3-mercaptophenylaniline (41.8mg, 0.3mmol) and 2-pyridinecarboxaldehyde (32.1mg, 0.3mmol) were employed to obtain 2-[2-(3-Methylmercaptophenyl)-2*H*-tetrazol-5-yl]pyridine as an orange solid.

$^1\text{H-NMR}$ (CDCl_3 , 300 MHz) δ 8.86 (d, 1H), 8.36 (d, 1H), 8.14 (t, 1H), 8.02 (d, 1H), 7.92 (t, 1H), 7.50-7.43 (m, 2H), 7.36 (d, 1H), 2.59 (s, 3H).

MS (ESI) 270.0 ($\text{M}^+ + \text{H}$).

15

EXAMPLE 15

2-[2-(4-fluorophenyl)-2*H*-tetrazol-5-yl]pyridine

Following the procedure described in **EXAMPLE 1** for the synthesis of 2-[2-(3-chlorophenyl)-2*H*-tetrazol-5-yl]pyridine, 4-fluorophenylaniline (33.3mg, 0.3mmol) and 2-pyridinecarboxaldehyde (32.1mg, 0.3mmol) were employed to obtain 2-[2-(4-fluorophenyl)-2*H*-tetrazol-5-yl]pyridine as an orange solid.

$^1\text{H-NMR}$ (CDCl_3 , 300 MHz) δ 8.85 (d, 1H), 8.35 (d, 1H), 8.28 (m, 2H), 7.92 (t, 1H), 7.49-7.44 (m, 1H), 7.29 (t, 2H).

MS (ESI) 242.1 ($\text{M}^+ + \text{H}$).

25

EXAMPLE 16

2-[2-(3-fluorophenyl)-2*H*-tetrazol-5-yl]pyridine

Following the procedure described in **EXAMPLE 1** for the synthesis of 2-[2-(3-chlorophenyl)-2*H*-tetrazol-5-yl]pyridine, 3-fluorophenylaniline (33.3mg, 0.3mmol) and 2-pyridinecarboxaldehyde (32.1mg, 0.3mmol) were employed to obtain 2-[2-(3-fluorophenyl)-2*H*-tetrazol-5-yl]pyridine as an orange solid.

$^1\text{H-NMR}$ (CDCl_3 , 300 MHz) δ 8.85 (d, 1H), 8.35 (d, 1H), 8.10 (d, 1H), 8.03 (d, 1H), 7.92 (t, 1H), 7.57 (q, 1H), 7.47 (m, 1H), 7.25 (t, 1H).

MS (ESI) 242.1 ($\text{M}^+ + \text{H}$).

EXAMPLE 17**2-[2-(2-methoxyphenyl)-2H-tetrazol-5-yl]pyridine**

Following the procedure described in **EXAMPLE 1** for the synthesis
5 of 2-[2-(3-chlorophenyl)-2H-tetrazol-5-yl]pyridine, 2-methoxyphenylaniline (36.9mg, 0.3mmol) and 2-pyridinecarboxaldehyde (32.1mg, 0.3mmol) were employed to obtain 2-[2-(2-methoxyphenyl)-2H-tetrazol-5-yl]pyridine as an orange solid.

¹H-NMR (CDCl₃, 300 MHz) δ 8.82 (d, 1H), 8.33 (d, 1H), 7.89 (t, 1H),
7.63 (d, 1H), 7.55 (t, 1H), 7.45-7.41 (m, 1H), 7.14 (d, 1H), 7.13 (t, 1H), 3.88 (s, 3H).

10 MS (ESI) 254.1 (M⁺+H).

EXAMPLE 18**2-[2-(3-ethylphenyl)-2H-tetrazol-5-yl]pyridine**

Following the procedure described in **EXAMPLE 1** for the synthesis
15 of 2-[2-(3-chlorophenyl)-2H-tetrazol-5-yl]pyridine, 3-ethylphenylaniline (36.4mg, 0.3mmol) and 2-pyridinecarboxaldehyde (32.1mg, 0.3mmol) were employed to obtain 2-[2-(3-ethylphenyl)-2H-tetrazol-5-yl]pyridine as an orange solid.

¹H-NMR (CDCl₃, 300 MHz) δ 8.84 (d, 1H), 8.36 (d, 1H), 8.13 (s, 1H),
8.08 (d, 1H), 7.91 (t, 1H), 7.51-7.42 (m, 2H), 7.35 (d, 1H), 2.78 (q, 2H), 1.32 (t, 3H).

20 MS (ESI) 252.2 (M⁺+H).

EXAMPLE 19**2-[2-(3-methylphenyl)-2H-tetrazol-5-yl]pyridine**

Following the procedure described in **EXAMPLE 1** for the synthesis
25 of 2-[2-(3-chlorophenyl)-2H-tetrazol-5-yl]pyridine, 3-methylphenylaniline (32.1mg, 0.3mmol) and 2-pyridinecarboxaldehyde (32.1mg, 0.3mmol) were employed to obtain 2-[2-(3-ethylphenyl)-2H-tetrazol-5-yl]pyridine as an orange solid.

¹H-NMR (CDCl₃, 300 MHz) δ 8.85 (d, 1H), 8.36 (d, 1H), 8.12 (s, 1H),
8.07 (d, 1H), 7.91 (t, 1H), 7.48-7.42 (m, 2H), 7.32 (d, 1H), 2.49 (s, 3H).

30 MS (ESI) 238.1 (M⁺+H).

EXAMPLE 20**2-[2-(2-chloro-3-pyridyl)-2H-tetrazol-5-yl]pyridine**

Following the procedure described in **EXAMPLE 1** for the synthesis
35 of 2-[2-(3-chlorophenyl)-2H-tetrazol-5-yl]pyridine, 3-amino-2-chloropyridine

(38.6mg, 0.3mmol) and 2-pyridinecarboxaldehyde (32.1mg, 0.3mmol) were employed to obtain 2-[2-(2-chloro-3-pyridyl)-2*H*-tetrazol-5-yl]pyridine as an orange solid.

¹H-NMR (CDCl₃, 300 MHz) δ 8.84 (d, 1H), 8.67 (d, 1H), 8.36 (d, 1H), 8.12 (d, 1H), 7.94 (t, 1H), 7.55 (m, 1H), 7.50-7.46 (m, 1H).

5 MS (ESI) 259.0 (M⁺+H).

EXAMPLE 21

2-[2-(3,5-dichlorophenyl)-2*H*-tetrazol-5-yl]pyridine

Following the procedure described in **EXAMPLE 1** for the synthesis
10 of 2-[2-(3-chlorophenyl)-2*H*-tetrazol-5-yl]pyridine, 3,5-dichloroaniline (48.6mg, 0.3mmol) and 2-pyridinecarboxaldehyde (32.1mg, 0.3mmol) were employed to obtain 2-[2-(3,5-dichlorophenyl)-2*H*-tetrazol-5-yl]pyridine as an orange solid.

¹H-NMR (CDCl₃, 300 MHz) δ 8.86 (d, 1H), 8.36 (d, 1H), 8.25 (d, 2H), 7.93 (t, 1H), 7.51-7.44 (m, 2H).

15 MS (ESI) 292.0 (M⁺+H).

EXAMPLE 22

2-[2-(2-chlorophenyl)-2*H*-tetrazol-5-yl]pyridine

Following the procedure described in **EXAMPLE 1** for the synthesis
20 of 2-[2-(3-chlorophenyl)-2*H*-tetrazol-5-yl]pyridine, 2-chloroaniline (38.27mg, 0.3mmol) and 2-pyridinecarboxaldehyde (32.1mg, 0.3mmol) were employed to obtain 2-[2-(2-chlorophenyl)-2*H*-tetrazol-5-yl]pyridine as an orange solid.

¹H-NMR (CDCl₃, 300 MHz) δ 8.84 (d, 1H), 8.34 (d, 1H), 7.92 (t, 1H), 7.71 (d, 1H), 7.66 (d, 1H), 7.59-7.43 (m, 3H).

25 MS (ESI) 258.0 (M⁺+H).

EXAMPLE 23

2-[2-(4-methoxyphenyl)-2*H*-tetrazol-5-yl]pyridine

Following the procedure described in **EXAMPLE 1** for the synthesis
30 of 2-[2-(3-chlorophenyl)-2*H*-tetrazol-5-yl]pyridine, 4-methoxyaniline (41.8mg, 0.3mmol) and 2-pyridinecarboxaldehyde (32.1mg, 0.3mmol) were employed to obtain 2-[2-(4-methoxyphenyl)-2*H*-tetrazol-5-yl]pyridine as an orange solid.

¹H-NMR (CDCl₃, 300 MHz) δ 8.84 (d, 1H), 8.34 (d, 1H), 8.19 (d, 2H), 7.90 (t, 1H), 7.46-7.42 (m, 1H), 7.07 (d, 2H), 3.90 (s, 3H).

35 MS (ESI) 254.1 (M⁺+H).

EXAMPLE 24**2-[2-(4-pyridyl)-2H-tetrazol-5-yl]pyridine**

Following the procedure described in **EXAMPLE 1** for the synthesis of 2-[2-(3-chlorophenyl)-2H-tetrazol-5-yl]pyridine, 4-aminopyridine (56.5mg, 0.6mmol) and 2-pyridinecarboxaldehyde (64.2mg, 0.6mmol) were employed to obtain 2-[2-(4-pyridyl)-2H-tetrazol-5-yl]pyridine as an orange solid.

¹H-NMR (CDCl₃, 300 MHz) δ 8.89 (d, 1H), 8.88 (d, 2H), 8.38 (d, 1H), 8.22 (d, 2H), 7.94 (t, 1H), 7.51-7.47 (m, 1H).

MS (ESI) 225.1 (M⁺+H).

EXAMPLE 25**2-[2-(3,5-dimethylphenyl)-2H-tetrazol-5-yl]pyridine**

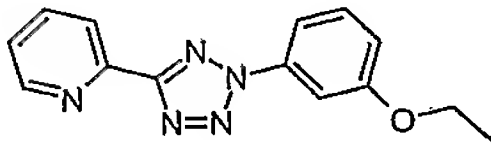
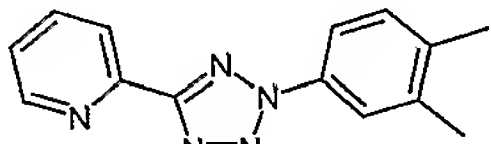
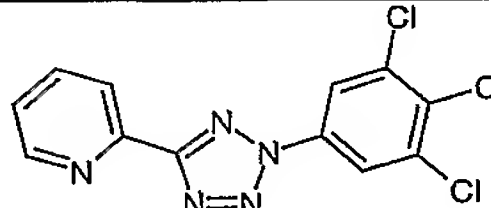
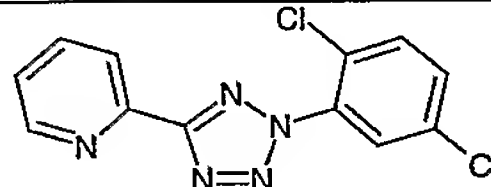
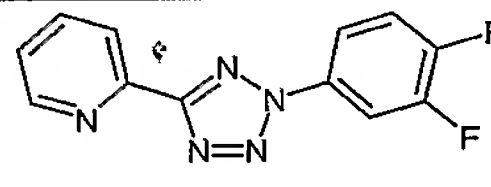
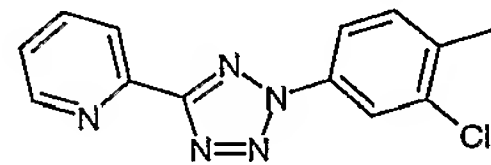
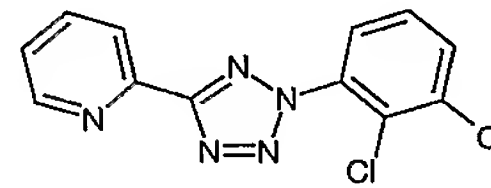
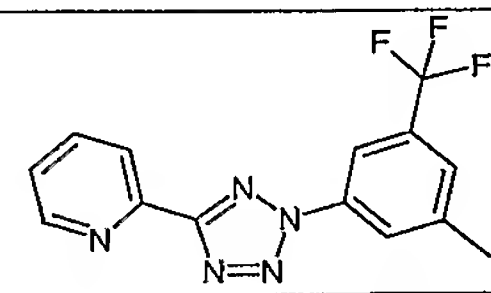
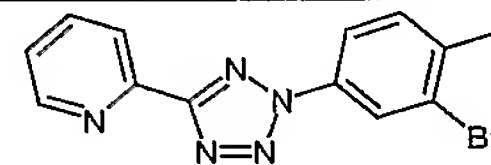
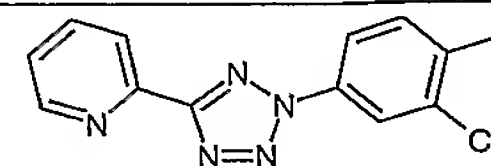
Following the procedure described in **EXAMPLE 1** for the synthesis of 2-[2-(3-chlorophenyl)-2H-tetrazol-5-yl]pyridine, 3,5-dimethylaniline (72.7mg, 0.6mmol) and 2-pyridinecarboxaldehyde (64.2mg, 0.6mmol) were employed to obtain 2-[2-(4-pyridyl)-2H-tetrazol-5-yl]pyridine as an orange solid. †

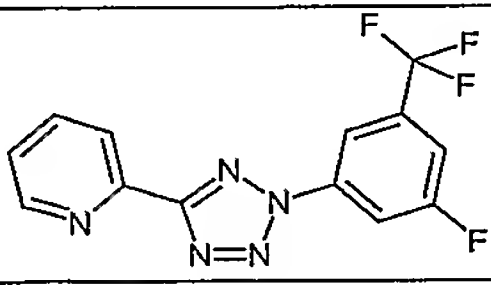
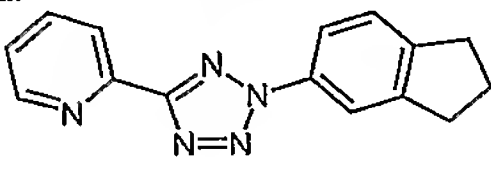
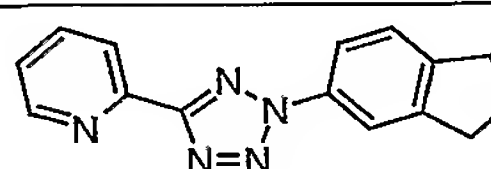
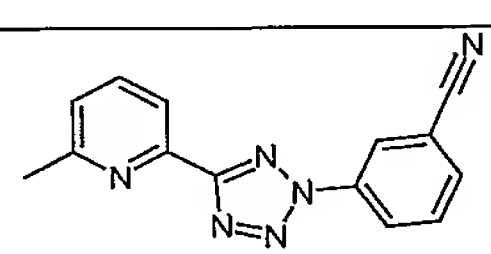
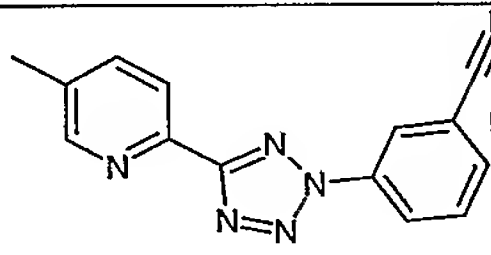
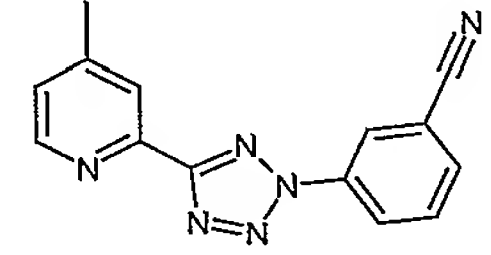
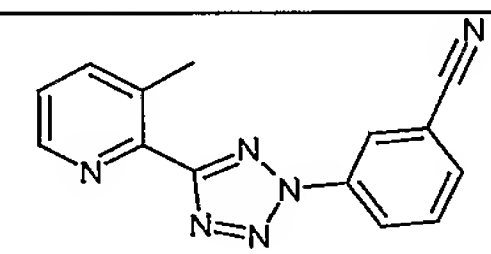
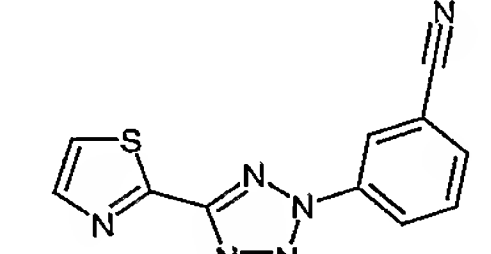
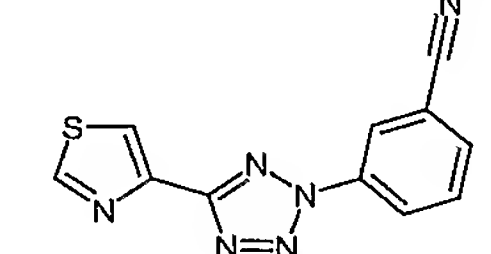
¹H-NMR (CDCl₃, 300 MHz) δ 8.84 (d, 1H), 8.35(d, 1H), 7.91 (s, 2H), 7.45-7.41 (m, 1H), 7.13 (s, 1H), 2.42 (s, 6H).

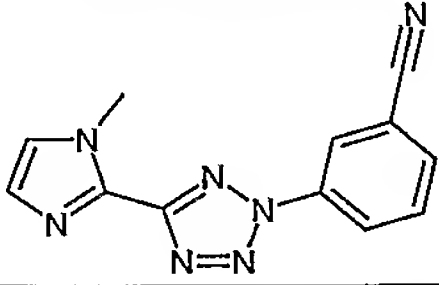
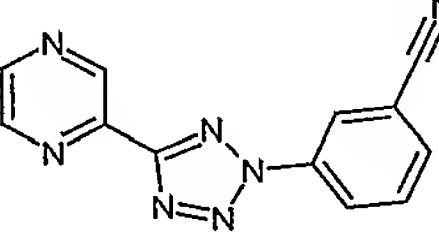
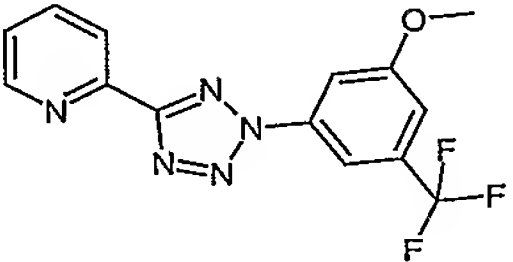
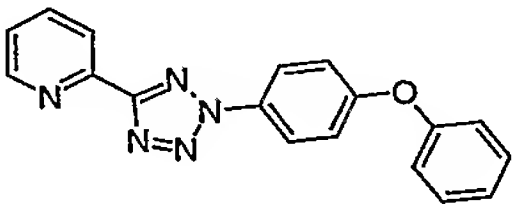
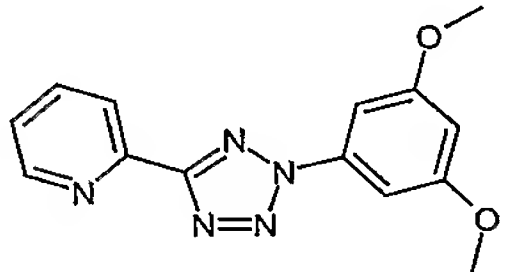
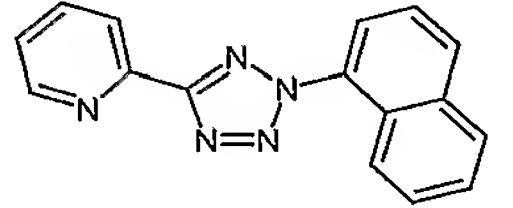
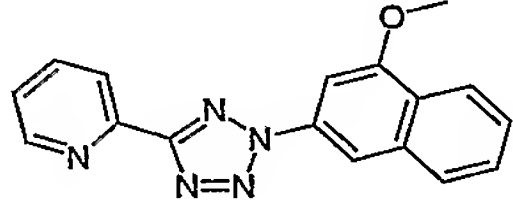
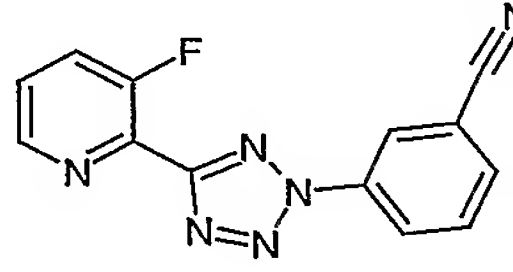
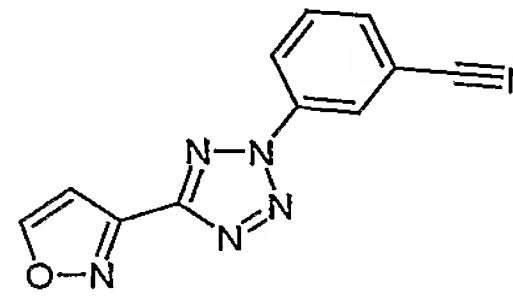
MS (ESI) 252.1 (M⁺+H).

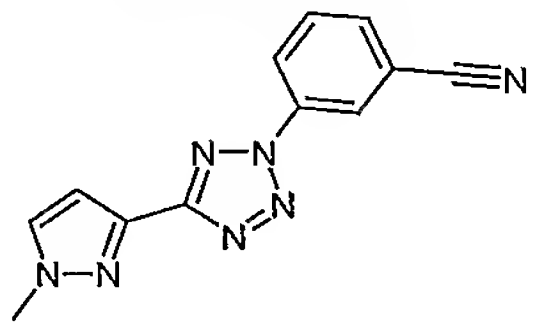
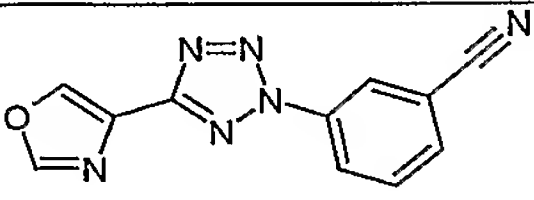
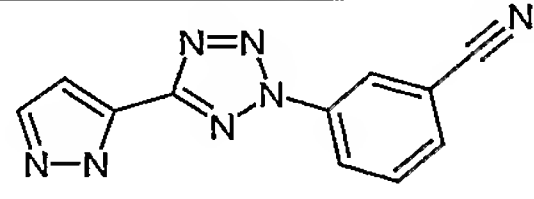
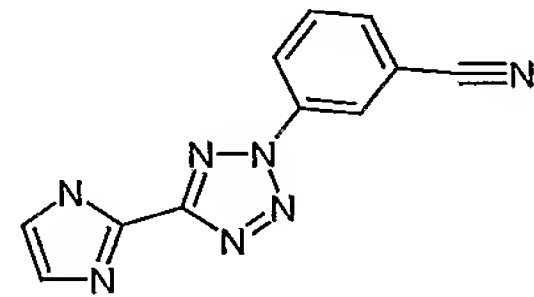
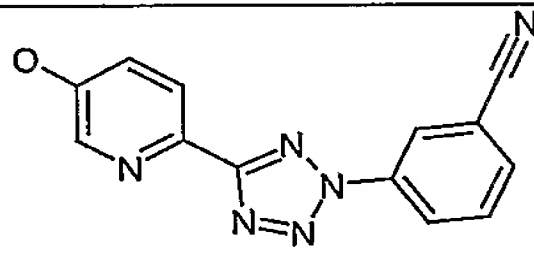
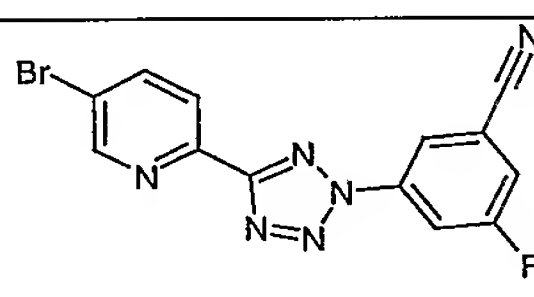
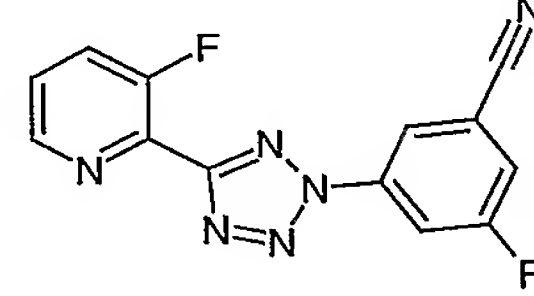
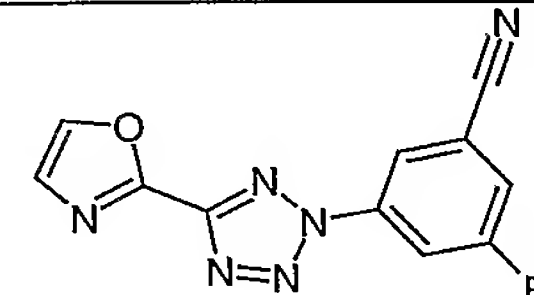
EXAMPLE 26 to EXAMPLE 206 shown below were prepared similarly to the schemes and procedures described above (ND = not determined).

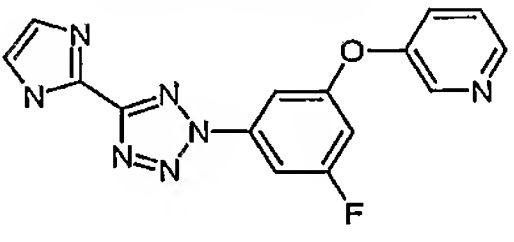
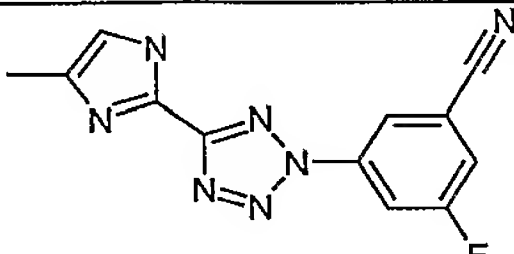
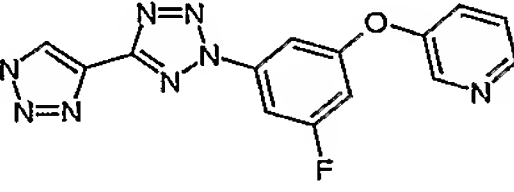
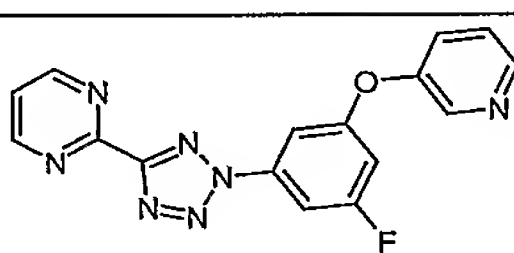
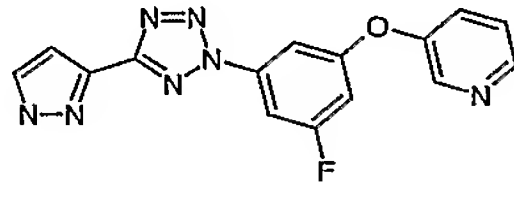
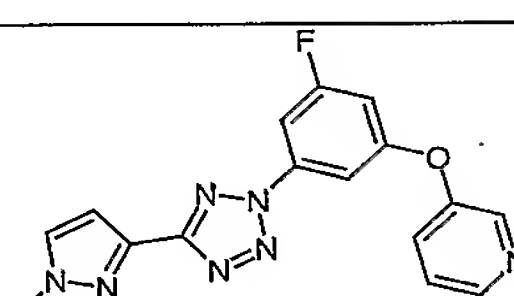
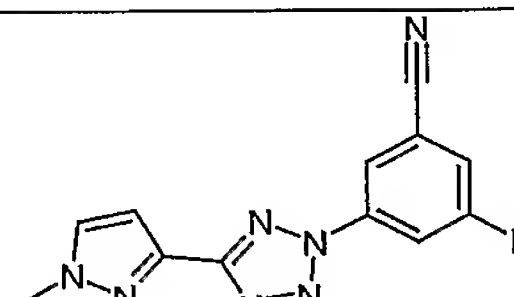
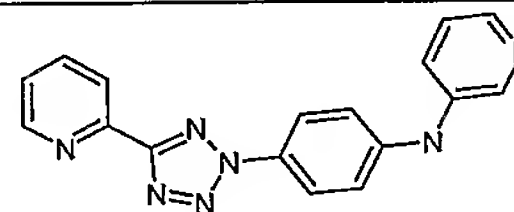
EXAMPLE	Structure	¹ H NMR	MS (ESI)
26		8.84-8.86 (d, 1H), 8.34-8.37 (d, 1H), 7.79-7.94 (m, 3H), 7.43-7.48 (m, 2H), 7.01-7.04 (m, 1H), 4.67-4.73 (m, 1H), 1.38-1.40 (d, 6H).	MS 282.1 (M ⁺ +H).
27		ND	MS 266.1 (M ⁺ +H).
28		ND	MS 308.0 (M ⁺ +H).

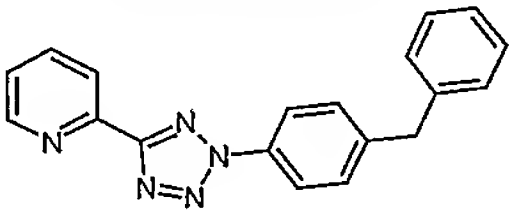
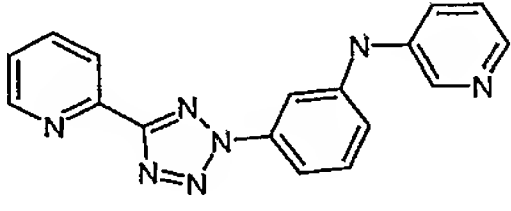
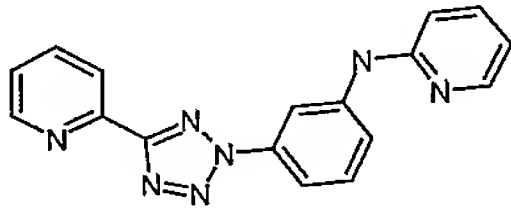
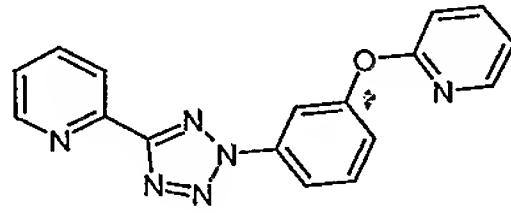
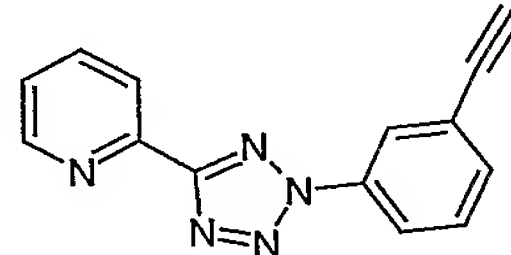
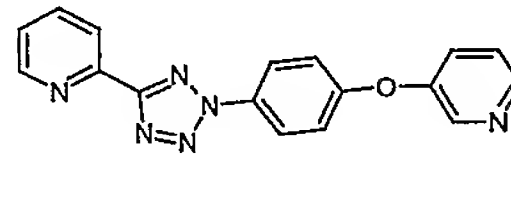
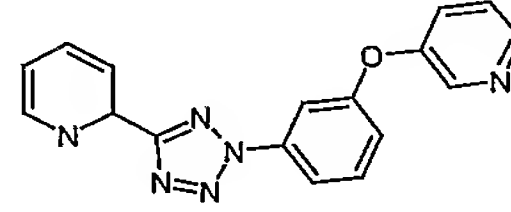
EXAMPLE	Structure	¹ H NMR	MS (ESI)
29		ND	MS 268.1 (M ⁺ +H).
30		8.83-8.85 (m, 1H), 8.34-8.37 (d, 1H), 8.06 (s, 1H), 7.90-8.00 (m, 2H), 7.43- 7.46 (m, 1H), 7.30-7.33 (d, 1H), 2.38 (s, 3H), 2.35 (s, 3H).	MS 252.1 (M ⁺ +H).
31		8.85-8.87 (d, 1H), 8.41 (s, 2H), 8.36- 8.38 (d, 1H), 7.91-7.97 (m, 1H), 7.46- 7.50 (m, 1H).	MS 326.0 (M ⁺).
32		8.83-8.85 (d, 1H), 8.34-8.36 (d, 1H), 7.90-7.96 (m, 1H), 7.79-7.80 (d, 1H), 7.45-7.62 (m, 3H).	MS 292.1 (M ⁺).
33		8.84-8.86 (d, 1H), 8.34-8.37 (d, 1H), 8.12-8.22 (m, 1H), 8.06-8.10 (m, 1H), 7.90-7.96 (m, 1H), 7.36-7.49 (m, 2H).	MS 260.2 (M ⁺ +H).
34		8.84-8.86 (d, 1H), 8.32-8.37 (m, 2H), 8.07-8.10 (m, 1H), 7.89-7.95 (m, 1H), 7.43-7.48 (m, 2H), 2.47 (s, 3H).	MS 272.2 (M ⁺ +H).
35		8.83-8.85 (d, 1H), 8.33-8.36 (d, 1H), 7.90-7.95 (m, 1H), 7.73-7.76 (m, 1H), 7.62-7.65 (m, 1H), 7.42-7.48 (m, 2H).	MS 292.1 (M ⁺).
36		8.85-8.87 (d, 1H), 8.36-8.39 (m, 2H), 8.34 (s, 1H), 7.90-7.96 (m, 1H), 7.59 (s, 1H), 7.45-7.49 (m, 1H), 2.57 (s, 3H).	MS 306.3 (M ⁺ +H).
37		8.84-8.86 (m, 1H), 8.50 (s, 1H), 8.34- 8.37 (m, 1H), 8.11-8.15 (m, 1H), 7.89- 7.94 (m, 1H), 7.42-7.48 (m, 2H), 2.49 (s, 3H).	MS 316.0 (M ⁺ +H).
38		ND	MS 384.0 (M ⁺).

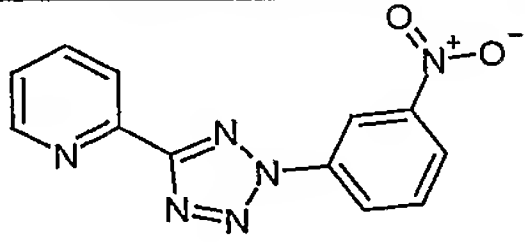
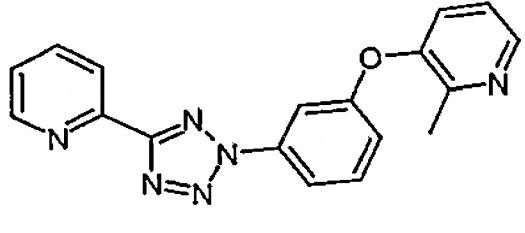
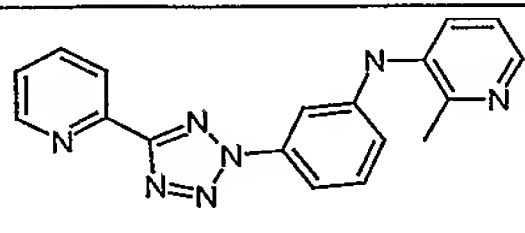
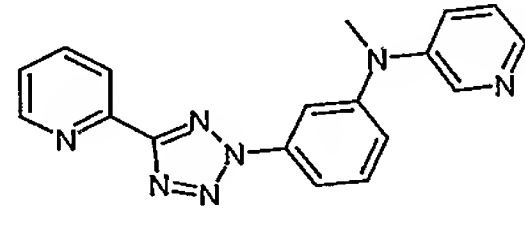
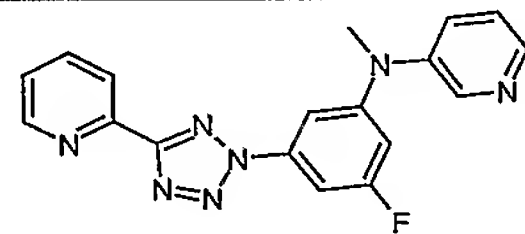
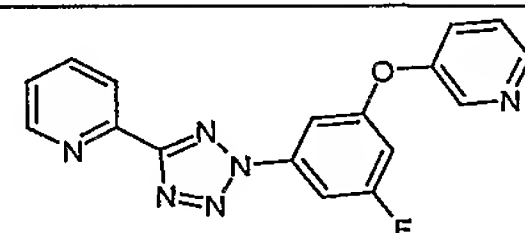
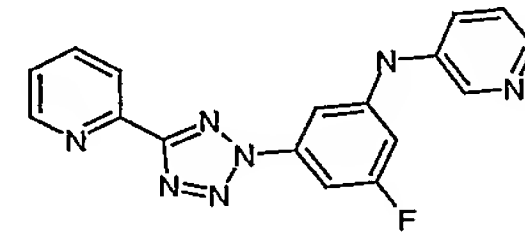
EXAMPLE	Structure	¹ H NMR	MS (ESI)
39		8.86-8.89 (m, 1H), 8.43 (s, 1H), 8.37-8.40 (d, 1H), 8.25-8.29 (m, 1H), 7.92-7.98 (m, 1H), 7.47-7.52 (m, 2H).	MS 310.0 (M ⁺ +H).
40		ND	MS 264.1 (M ⁺ +H).
41		8.84-8.86 (m, 1H), 8.35-8.37 (m, 1H), 8.19-8.23 (m, 1H), 8.17 (s, 1H), 7.89-7.95 (m, 1H), 7.43-7.48 (m, 2H), 5.21 (s, 4H).	MS 266.0 (M ⁺ +H).
42		8.75-8.82 (m, 2H), 8.60-8.65 (m, 2H), 8.20-8.22 (d, 1H), 8.02-8.05 (d, 1H), 7.92-7.97 (m, 1H), 3.06 (s, 1H).	MS 263.1 (M ⁺ +H).
43		8.80 (s, 1H), 8.63-8.66 (d, 1H), 8.57-8.58 (m, 1H), 8.49-8.54 (m, 2H), 7.93-7.95 (d, 1H), 7.81-7.85 (dd, 1H), 2.58 (s, 1H).	MS 263.1 (M ⁺ +H).
44		9.09-9.12 (d, 1H), 9.00 (s, 1H), 8.90 (s, 1H), 8.82-8.86 (m, 1H), 8.32-8.34 (d, 1H), 8.26-8.29 (d, 1H), 8.13-8.18 (dd, 1H), 3.06 (s, 1H).	MS 263.1 (M ⁺ +H).
45		8.65-8.70 (d, 1H), 8.47-8.52 (m, 2H), 8.39-8.43 (d, 1H), 7.91-7.97 (dd, 1H), 7.83-7.85 (d, 1H), 7.69-7.74 (dd, 1H), 2.80 (s, 1H).	MS 263.1 (M ⁺ +H).
46		8.61 (s, 1H), 8.53-8.56 (d, 1H), 8.12-8.13 (d, 1H), 7.83-7.86 (d, 1H), 7.74-7.79 (dd, 1H), 7.64-7.65 (d, 1H).	MS 255.1 (M ⁺ +H).
47		8.61 (s, 1H), 8.53-8.56 (d, 1H), 8.12-8.13 (d, 1H), 7.83-7.86 (d, 1H), 7.74-7.79 (dd, 1H), 7.65-7.66 (d, 1H).	MS 255.1 (M ⁺ +H).

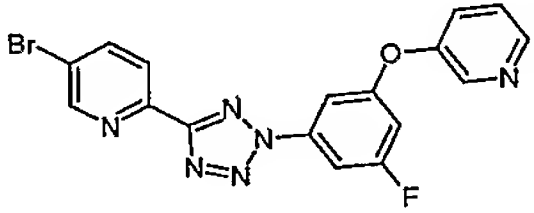
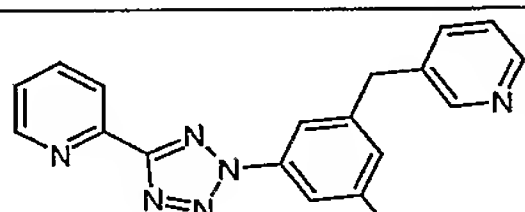
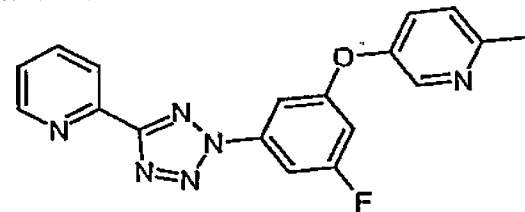
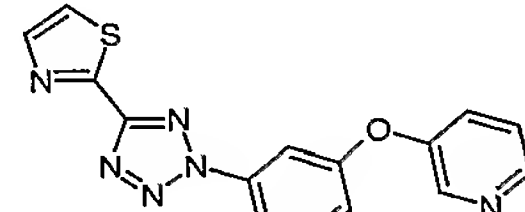
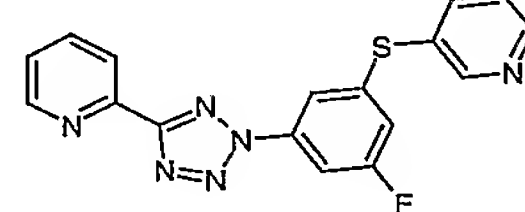

EXAMPLE	Structure	¹ H NMR	MS (ESI)
48		8.58-8.59 (m, 1H), 8.51-8.55 (m, 1H), 7.81-7.84 (d, 1H), 7.72-7.77 (dd, 1H), 7.28 (s, 1H), 7.13 (s, 1H), 4.19 (s, 3H).	MS 252.1 (M ⁺ +H).
49		9.55-9.57 (s, 1H), 8.81-8.87 (m, 2H), 8.67 (s, 1H), 8.57-8.62 (m, 1H), 7.94-7.97 (d, 1H), 7.84-7.91 (dd, 1H).	MS 250.1 (M ⁺ +H).
50		8.85-8.87 (d, 1H), 8.36-8.39 (d, 1H), 8.16 (s, 1H), 8.00-8.01 (t, 1H), 7.90-7.96 (t, 1H), 7.45-7.49 (m, 1H), 7.29 (s, 1H).	MS 322.0 (M ⁺ +H).
51		8.83-8.85 (d, 1H), 8.33-8.36 (d, 1H), 8.20-8.23 (d, 2H), 7.87-7.93 (t, 1H), 7.39-7.46 (m, 3H), 7.08-7.23 (m, 5H).	MS 316.1 (M ⁺ +H).
52		(HCl salt in CD ₃ OD) 8.95-9.05 (br, 1H), 8.84-8.87 (d, 1H), 8.76-8.80 (t, 1H), 8.19-8.25 (t, 1H), 7.40-7.41 (d, 2H), 6.74-6.76 (t, 1H), 3.92 (s, 6H).	MS 284.6 (M ⁺ +H).
53		8.85-8.87 (d, 1H), 8.37-8.40 (d, 1H), 8.07-8.11 (m, 2H), 7.90-8.00 (m, 3H), 7.59-7.67 (m, 3H), 7.43-7.48 (m, 1H).	MS 274.6 (M ⁺ +H).
54		ND	MS 304.6 (M ⁺ +H).
55		8.71 – 8.72 (m, 1H), 8.66 (m, 1H), 8.51 – 8.54 (m, 1H), 8.15 – 8.16 (m, 1H), 8.06 – 8.10 (m, 1H), 7.94 (dd, 1H), 7.77 – 7.80 (m, 1H).	MS 267.4 (M ⁺ +H).
56		9.32 (d, 1H), 8.65 – 8.67 (m, 1H), 8.50 – 8.52 (m, 1H), 8.14 – 8.15 (m, 1H), 7.91 (d, 1H), 7.32 (d, 1H).	MS 239.1 (M ⁺ +H).

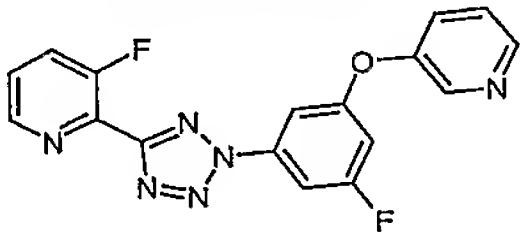
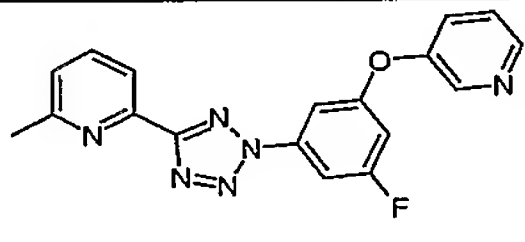
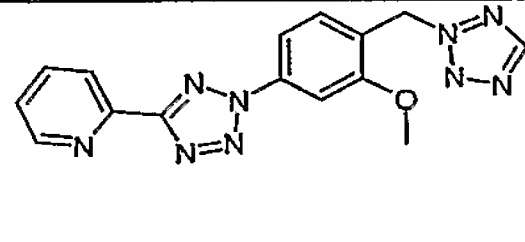
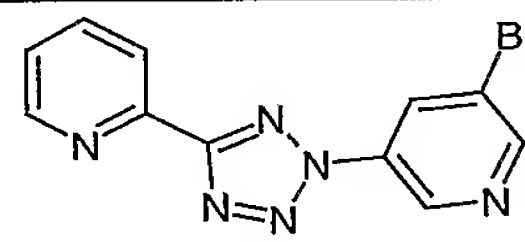
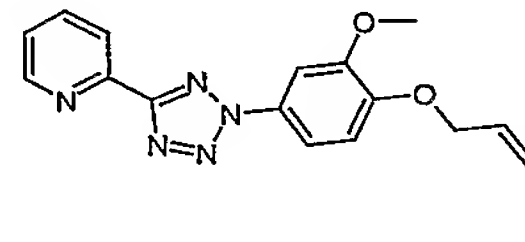
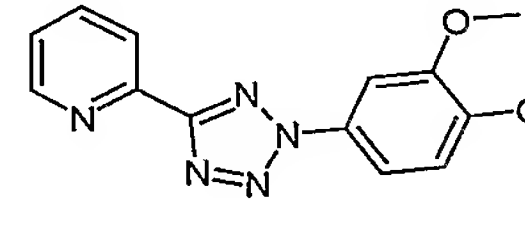
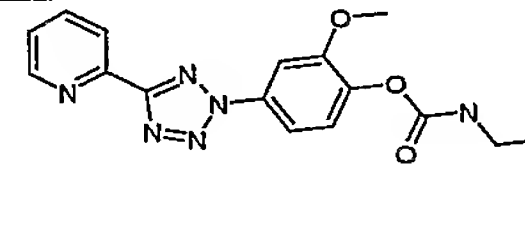
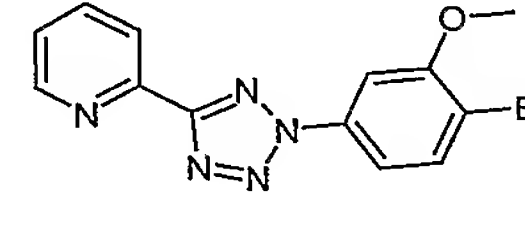
EXAMPLE	Structure	¹ H NMR	MS (ESI)
57		8.59 – 8.60 (m, 1H), 8.45 – 8.48 (m, 1H), 8.08 – 8.10 (m, 1H), 7.94 (d, 1H), 7.87 (dd, 1H), 6.91 (d, 1H), 3.98, (s, 3H).	MS 252.1 (M ⁺ +H).
58		8.80 (d, 1H), 8.63 – 8.64 (m, 1H), 8.55 – 8.57 (m, 1H), 8.51 (d, 1H), 7.97 – 7.99 (m, 1H), 7.87 (dd, 1H).	MS 239.0 (M ⁺ +H).
59		8.61 (m, 1H), 8.44 – 8.50 (m, 1H), 8.11 – 8.12 (m, 1H), 8.00 (m, 1H), 7.91 (dd, 1H), 6.96 (m, 1H).	MS 238.0 (M ⁺ +H).
60		8.68 – 8.69 (m, 1H), 8.60 – 8.62 (m, 1H), 8.06 – 8.09 (m, 1H), 7.94 (dd, 1H), 7.87 (s, 2H).	MS 238.0 (M ⁺ +H).
61		10.51 (s, 1H), 8.66 (m, 1H), 8.50 – 8.52 (m, 1H), 8.27 – 8.28 (m, 1H), 8.12 – 8.14 (m, 1H), 7.93 (dd, 1H), 7.47 – 7.53 (m, 2H).	MS 265.1 (M ⁺ +H).
62		8.96 (m, 1H), 8.57 (m, 1H), 8.45 – 8.47 (m, 1H), 8.36 – 8.38 (m, 1H), 8.21 – 8.24 (m, 2H).	MS 345.2 (M ⁺ +H).
63		8.71 – 8.73 (m, 1H), 8.56 (m, 1H), 8.45 – 8.47 (m, 1H), 8.22 – 8.25 (m, 1H), 8.07 – 8.10 (m, 1H), 7.77 – 7.81 (m, 1H).	MS 285.0 (M ⁺ +H).
64		8.45 (m, 1H), 8.27 – 8.36 (m, 1H), 7.98 (s, 1H), 7.58 – 7.60 (m, 1H), 7.49 (m, 1H).	MS 257.0 (M ⁺ +H).

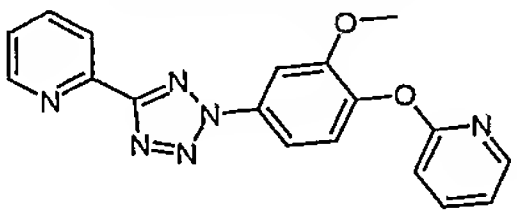
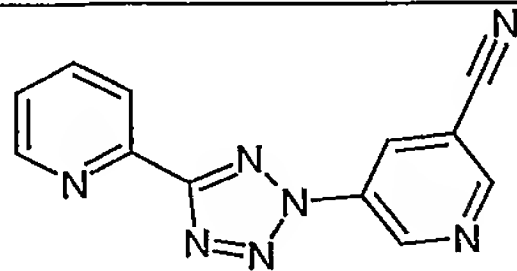
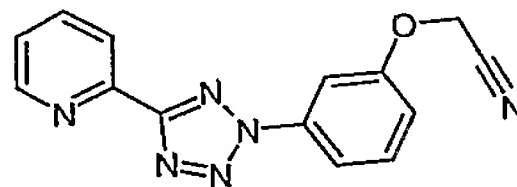
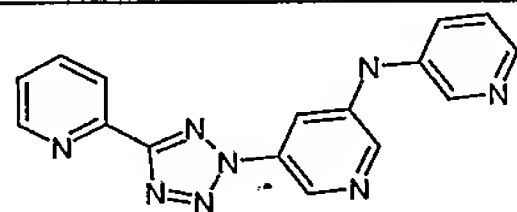
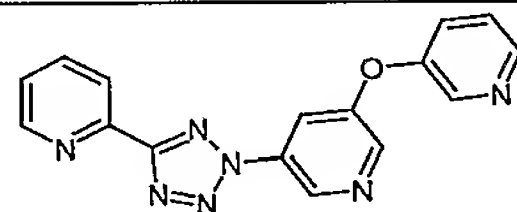
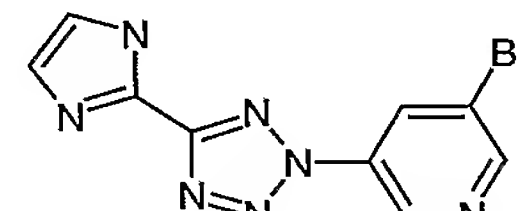
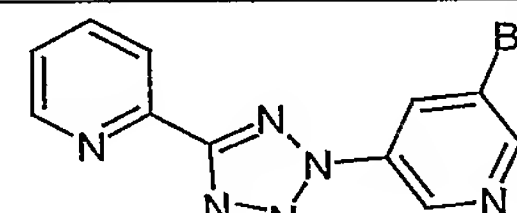
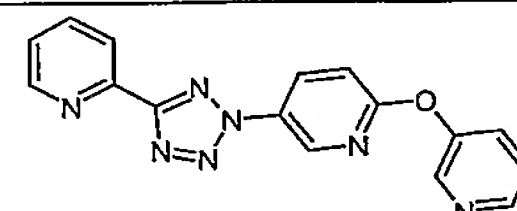
EXAMPLE	Structure	¹ H NMR	MS (ESI)
65		8.91 (m, 1H), 8.74 (m, 1H), 8.38 – 8.40 (m, 1H), 8.09 (dd, 1H), 8.04 – 8.07 (m, 2H), 7.91 (s, 2H), 7.41 – 7.43 (m, 1H)	MS 324.3 (M ⁺ +H).
66		8.51 (m, 1H), 8.40 – 8.42 (m, 1H), 8.23 – 8.25 (m, 1H), 7.23 (m, 1H), 2.30 (s, 3H).	MS 270.5 (M ⁺ +H).
67		8.61 (d, 1H), 8.53 – 8.54 (m, 1H), 7.85 – 7.87 (m, 1H), 7.75 – 7.78 (m, 1H), 7.60 (d, 1H), 7.54 (dd, 1H), 7.33 – 7.36 (m, 1H).	MS 325.1 (M ⁺ +H).
68		9.09 (d, 2H), 8.67 – 8.68 (m, 1H), 8.53 – 8.62 (m, 1H), 7.90 – 7.92 (m, 1H), 7.86 – 7.88 (m, 1H), 7.72 (t, 1H), 7.61 – 7.66 (m, 2H), 7.39 – 7.42 (m, 1H).	MS 336.3 (M ⁺ +H).
69		8.65 (d, 1H), 8.56 – 8.57 (m, 1H), 7.97 (d, 1H), 7.83 – 7.87 (m, 2H), 7.60 – 7.64 (m, 2H), 7.34 – 7.36 (m, 1H), 6.93 (d, 1H).	MS 324.0 (M ⁺ +H).
70		8.70 (d, 1H), 8.58 – 8.60 (m, 1H), 7.94 (d, 1H), 7.90 – 7.92 (m, 1H), 7.86 – 7.88 (m, 1H), 7.68 (dd, 1H), 7.64 (m, 1H), 7.36 – 7.38 (m, 1H), 6.90 (d, 1H), 4.02 (s, 3H).	MS 338.3 (M ⁺ +H).
71		8.51 (s, 1H), 8.40 – 8.43 (m, 1H), 8.13 – 8.19 (m, 1H), 7.96 (d, 1H), 6.94 (d, 1H), 4.00 (s, 3H).	MS 270.3 (M ⁺ +H).
72		8.9-8.85 (br, 1H), 8.57-8.56 (m, 2H), 8.41-8.37 (m, 1H), 8.30-8.27 (m, 4H), 7.9 (dd, 1H), 7.87 (br, 1H), 7.57 (d, 2H).	MS 315.98 (M ⁺ +H).

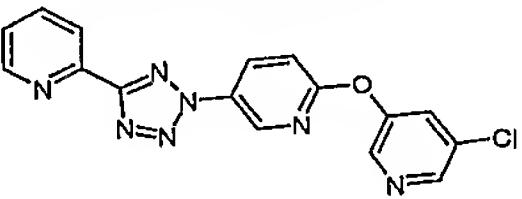
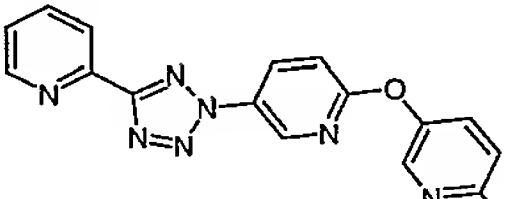
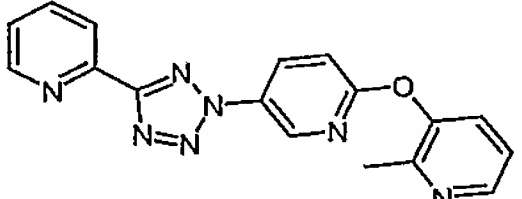
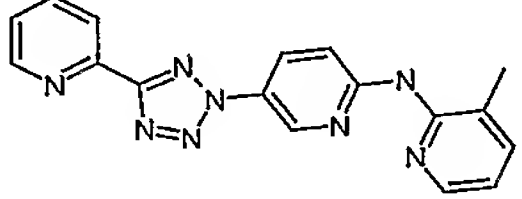
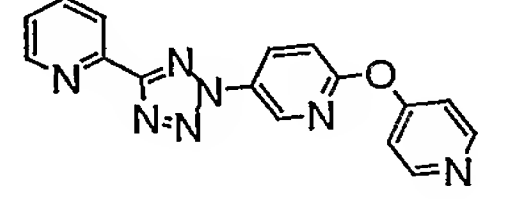
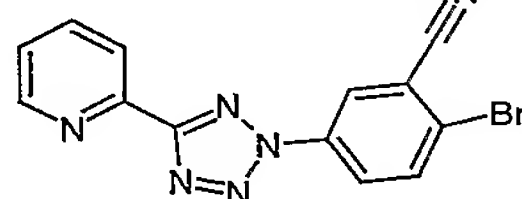
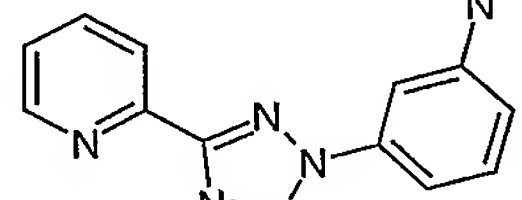
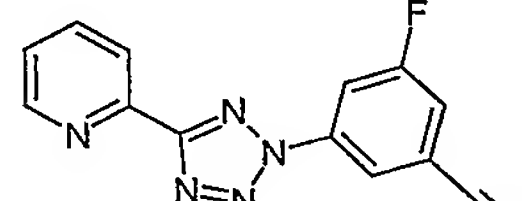
EXAMPLE	Structure	¹ H NMR	MS (ESI)
73		8.96 (br, 1H), 8.78 (br, 1H) 8.69-8.64 (dd, 1H), 8.22-8.19 (m, 2H), 8.11-8.107 (m, 1H), 7.57-7.55 (m, 2H), 7.35-7.24 (m, 5H), 4.15 (s, 2H).	MS 314.77 (M ⁺).
74		8.91 (br, 1H), 8.66-8.646 (m, 1H), 8.57-8.56 (m, 1H), 8.50-8.47 (m, 1H), 8.30-8.27 (m, 2H), 8.18-8.17 (m, 1H), 8.03-8.02 (m, 1H), 7.95-7.91 (m, 2H), 7.74-7.71 (m, 1H), 7.56-7.54 (m, 1H).	MS 316.03 (M ⁺ +H).
75		8.97 (br, 1H) 8.78-8.76 (m, 1H), 8.65 (t, 1H), 8.36 (s, 1H), 8.32-8.31 (d, 1H), 8.18 (t, 1H), 8.09 (br, 1H), 8.03-8.02 (d, 1H), 7.88 (t, 1H), 7.71-7.70 (d, 1H), 7.41-7.40 (d, 1H), 7.19 (t, 1H).	MS 316.09 (M ⁺ +H).
76		8.90 (br, 1H), 8.67 (br, 1H), 8.48 (t, 1H), 8.23-8.22 (m, 1H), 8.17-8.15 (dd, 1H), 8.07 (t, 1H), 8.00-7.96 (m, 2H), 7.77-7.73 (t, 1H), 7.43-7.41 (dd, 1H), 7.27-7.24 (dd, 1H), 7.18-7.16 (d, 1H).	MS 317.03 (M ⁺ +H).
77		9.00 (br, 1H), 8.83 (br, 1H), 8.74-8.71 (t, 1H), 8.35 (s, 1H), 8.30-8.29 (d, 1H), 8.16 (br, 1H), 7.76-7.69 (m, 2H), 3.81 (s, 1H).	MS 248.06 (M ⁺ +H).
78		9.02 (br, 1H), 8.90-8.88 (m, 1H), 8.84-8.80 (t, 1H), 8.45-8.41 (m, 3H), 8.26-8.19 (t, 2H), 7.64-7.62 (d, 2H), 7.50-7.48 (m, 1H), 7.30-7.28 (m, 1H).	MS 316.49 (M ⁺).
79		8.94 (br, 1H), 8.89 (br, 1H), 8.71 (br, 1H), 8.69 (br, 1H), 8.52 (t, 1H), 8.41-8.39 (m, 1H), 8.28-8.26 (d, 1H), 8.20 (br, 1H), 8.15-8.12 (m, 1H), 8.00 (br, 1H), 7.85 (t, 1H), 7.55-7.53 (d, 1H).	MS 317.05 (M ⁺ +H).

EXAMPLE	Structure	¹ H NMR	MS (ESI)
80		9.18 (t, 1H), 8.88 (d, 1H), 8.67 (ddd, 1H), 8.41-8.38 (m, 2H), 7.97-8.38 (m, 1H), 7.83 (t, 1H), 7.49 (ddd, 1H).	MS 269.05 (M ⁺ +H).
81		8.84 (d, 1H), 8.39 (dd, 1H), 8.34 (d, 1H), 8.04-8.02 (m, 1H), 7.94-7.90 (m, 1H), 7.88-7.87 (m, 1H), 7.54 (t, 1H), 7.46-7.44 (m, 1H), 7.30 (dd, 1H), 7.19 (dd, 1H), 7.06-7.04 (m, 1H), 2.54 (s, 3H).	MS 331.17 (M ⁺ +H).
82		8.84 (ddd, 1H), 8.35-8.33 (m, 1H), 8.27 (dd, 1H), 7.92-7.89 (m, 1H), 7.83-7.80 (m, 2H), 7.61 (dd, 1H), 7.46-7.43 (t, 2H), 7.16 (dd, 1H), 7.04-7.03 (m, 1H), 2.55 (s, 3H).	MS 330.13 (M ⁺ +H).
83		8.76-8.75 (d, 1H), 8.37-8.34 (m, 2H), 8.24-8.20 (m, 3H), 8.10-8.07 (m, 1H), 8.03-8.01 (dd, 1H), 7.86-7.80 (m, 2H), 7.63-7.59 (m, 2H), 3.56 (s, 3H).	MS 352.25 (M ⁺ +Na).
84		8.78 (br, 1H), 8.50 (br, 1H), 8.38 (br, 1H), 8.319 (br, 1H), 8.18-8.15 (dd, 1H), 8.11-8.09 (t, 1H), 8.06 (s, 1H), 7.97-7.95 (m, 1H), 7.92-7.89 (m, 1H), 7.64 (br, 1H), 7.42-7.39 (m, 1H), 3.57 (s, 3H).	MS 348.22 (M ⁺ +H).
85		8.84-8.82 (ddd, 1H), 8.51-8.50 (m, 2H), 8.34-8.32 (m, 1H), 7.92-7.89 (m, 1H), 7.83-7.81 (m, 1H), 7.76-7.75 (m, 1H), 7.46-7.43 (m, 2H), 7.39-7.36 (m, 1H), 6.87-6.85 (m, 1H).	MS 335.28 (M ⁺ +H).
86		8.87-8.85 (m, 1H), 8.544-8.540 (m, 1H), 8.39-8.35 (m, 2H), 7.95-7.92 (m, 1H), 7.741-7.737 (m, 1H), 7.62-7.55 (m, 2H), 7.49-7.46 (m, 1H), 7.35-7.32 (dd, 1H),	MS 334.21 (M ⁺ +H).

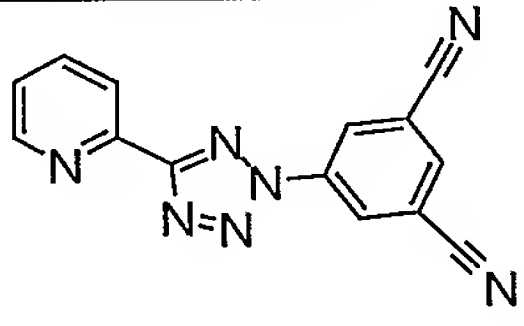
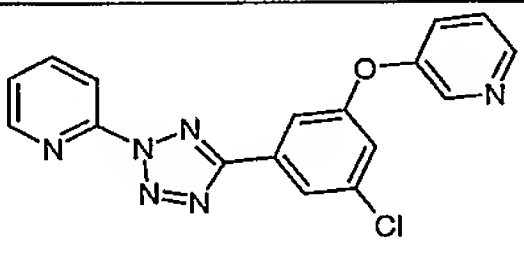
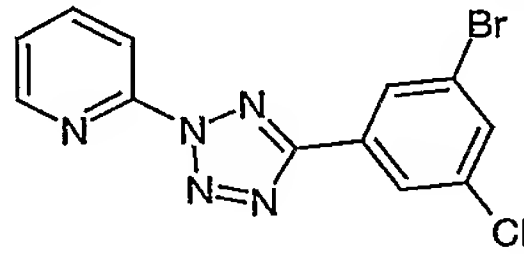
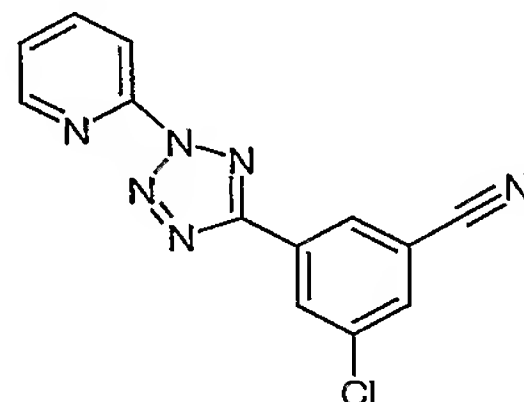
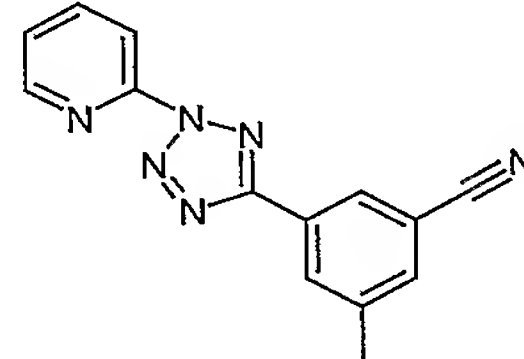
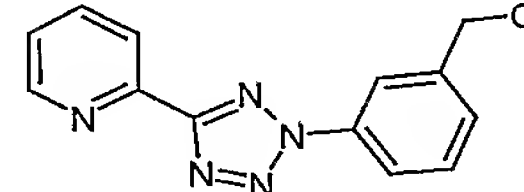
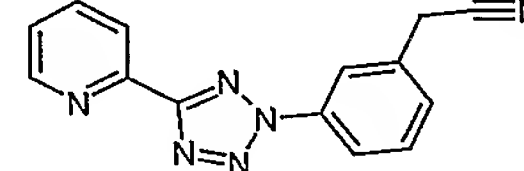
EXAMPLE	Structure	¹ H NMR	MS (ESI)
		6.89-6.86 (m 1H), 6.26 (s, 1H).	
87		8.889-8.885 (m, 1H), 8.53-8.52 (m, 2H), 8.24-8.22 (d, 1H), 8.06-8.04 (dd, 1H), 7.82-7.80 (m, 1H), 7.74 (s, 1H), 7.46-7.44 (m, 1H), 7.40-7.38 (dd, 1H), 6.87-6.85 (m 1H).	MS 386.91 (M ⁺ +H-N ₂).
88		8.86-8.85 (m, 1H), 8.56 (br, 1H), 8.535-8.527 (d, 1H), 8.36-8.35 (d, 1H), 8.00 (s, 1H), 7.94-7.90 (m, 2H), 7.54-7.51 (m, 1H), 7.47-7.44 (m, 1H), 7.283-7.258 (m, 1H), 7.03-7.02 (m, 1H), 4.10 (s, 2H).	MS 332.94 (M ⁺ +H).
89		8.85-8.83 (m, 1H), 8.394-8.389 (d, 1H), 8.34-8.32 (m, 1H), 7.93-7.90 (m, 1H), 7.79-7.77 (m, 1H), 7.72-7.71 (m, 1H), 7.47-7.44 (ddd, 1H), 7.37-7.28 (dd, 1H), 7.24-7.23 (d, 1H), 6.84-6.81 (m, 1H), 2.61 (s, 3H).	MS 349.23 (M ⁺ +H).
90		8.915-8.909 (d, 1H), 8.73-8.72 (d, 1H), 8.45-8.42 (ddd, 1H), 8.14-8.11 (m, 2H), 8.04-8.01 (m, 1H), 7.99-7.98 (m, 1H), 7.95 (d, 1H), 7.38-7.35 (dd, 1H).	MS 341.21 (M ⁺ +H).
91		8.85 (dd, 1H), 8.74 (d, 1H), 8.65 (dd, 1H), 8.34-8.33 (m, 1H), 8.06 (s, 1H), 7.93-7.87 (m, 2H), 7.85-7.83 (m, 1H), 7.48-7.45 (m, 1H), 7.39-7.36 (m, 1H), 7.00-6.98 (m, 1H).	MS 350.86 (M ⁺ +H).
92		8.86-8.85 (m, 1H), 8.432-8.428 (m, 1H), 8.38-8.36 (d, 1H), 8.296-8.289 (m, 1H), 8.20 (s, 1H), 8.03-8.00 (m, 1H), 7.95-7.91 (m, 1H), 7.48-7.45 (m, 1H), 7.35-7.33 (m, 1H), 7.31-7.26 (m, 2H), 5.22 (s, 2H).	MS 349.36 (M ⁺ +H).

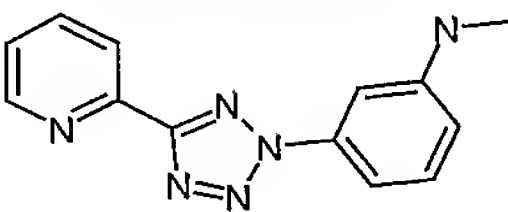
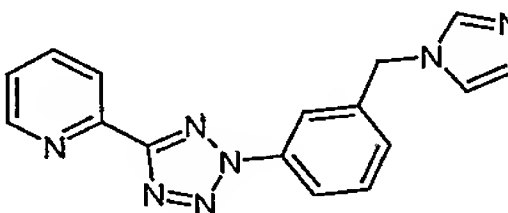
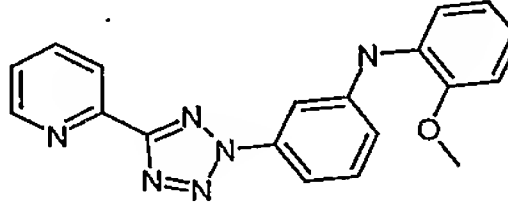
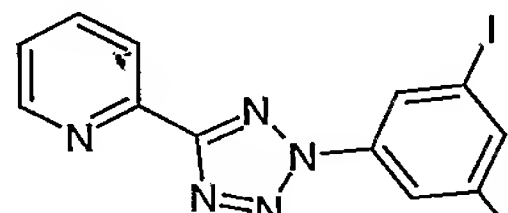
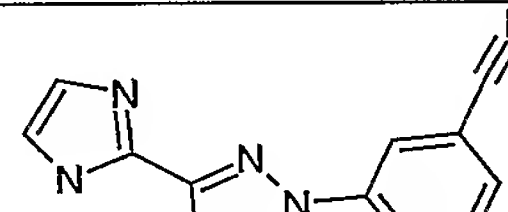
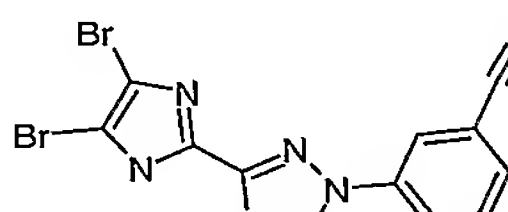

EXAMPLE	Structure	¹ H NMR	MS (ESI)
93		8.69-8.68 (d, 1H), 8.52 (br, 2H), 7.83-7.80 (m, 1H), 7.77 (m, 2H), 7.70-7.66 (m, 1H), 7.54-7.50 (m, 1H), 7.46-7.45 (m, 1H), 7.40-7.38 (m, 1H), 6.87-6.84 (m, 1H).	MS 353.30 (M ⁺ +H).
94		8.94 (d, 1H), 8.75-8.74 (d, 1H), 8.59-8.58 (d, 1H), 8.53 (t, 1H), 8.46 (ddd, 1H), 8.16 (dd, 1H), 8.09-8.07 (m, 2H), 7.98 (d, 1H), 7.43-7.41 (m, 1H), 2.91 (s, 3H).	MS 348.99 (M ⁺ +H).
95		8.97-8.94 (d, 1H), 8.85-8.87 (d, 1H), 8.78 (s, 1H), 8.67-8.64 (m, 1H), 8.11-8.08 (t, 1H), 7.95-7.91 (m, 2H), 7.58-7.57 (d, 1H), 6.02 (s, 2H), 4.03 (s, 3H).	MS 336.3 (M ⁺ +H).
96		9.48 (s, 1H), 9.07 (m, 1H), 8.99 (s, 1H), 8.89-8.95 (m, 2H), 8.86 (dd, 1H), 8.28 (dd, 1H).	MS 302.97 (M ⁺ +H).
97		8.85 (s, 1H), 8.36 (d, 1H), 7.79-7.73 (m, 3H), 7.45 (m, 1H), 7.01 (d, 1H), 6.15 (m, 1H), 5.46 (d, 1H), 5.32 (d, 1H), 4.70 (m, 2H), 4.03 (s, 3H).	MS 310.65 (M ⁺ +H).
98		8.95 (d, 1H), 8.78 (d, 1H), 8.69 (m, 1H), 8.11 (m, 1H), 7.72-7.78 (m, 2H), 7.06 (m, 1H), 4.03 (s, 3H).	MS 270.40 (M ⁺ +H).
99		8.80 (s, 1H), 8.73 (m, 1H), 8.62 (m, 1H), 7.83-8.07 (m, 3H), 7.36 (d, 1H), 3.99 (s, 3H), 3.22 (q, 2H), 1.14 (t, 3H).	MS 340.99 (M ⁺ +H).
100		8.97 (d, 1H), 8.79 (d, 1H), 8.69 (dd, 1H), 8.12 (dd, 1H), 7.78-7.91 (m, 3H), 4.06 (s, 3H).	MS 332.14 (M ⁺ +H).

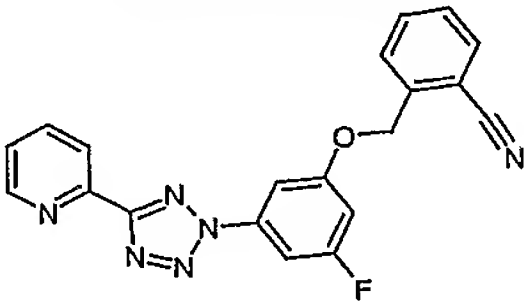
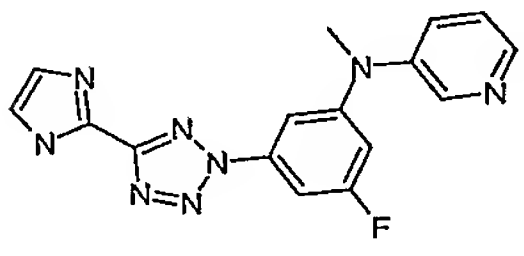
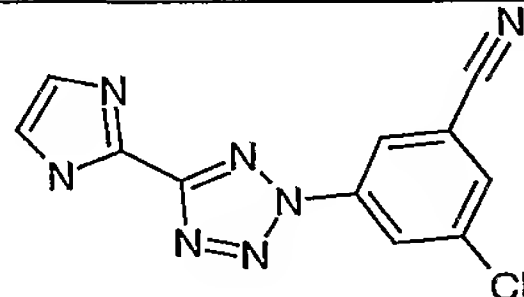
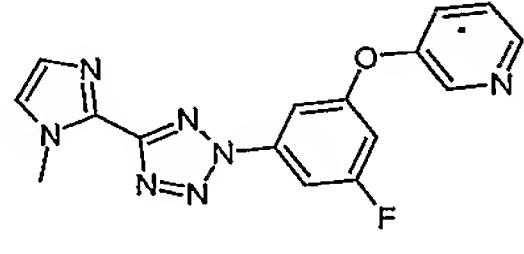
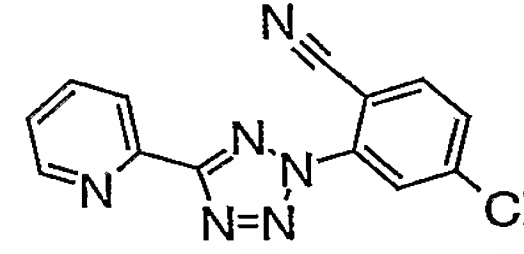
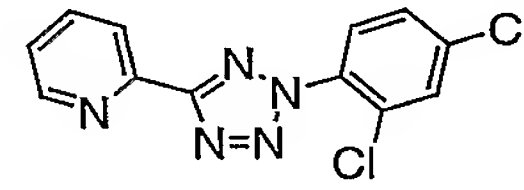
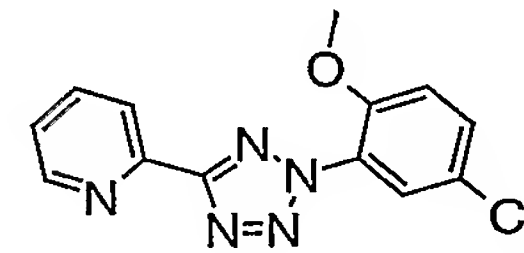
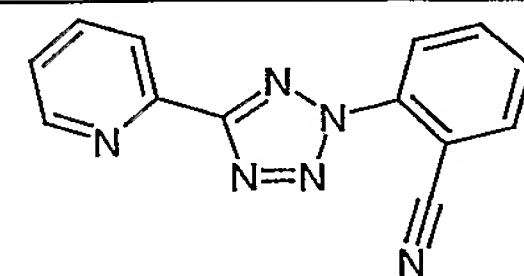
EXAMPLE	Structure	¹ H NMR	MS (ESI)
101		9.04 (d, 1H), 8.91 (d, 1H), 8.27 (dd, 1H), 8.47 (d, 1H), 8.37 (dd, 1H), 8.25 (dd, 1H), 8.13 (s, 1H), 8.04 (m, 1H), 7.70 (d, 1H), 7.58 (dd, 1H), 7.28 (d, 1H), 3.98 (s, 3H).	MS 347.44 (M ⁺ +H).
102		9.76 (d, 1H), 9.08 (d, 1H), 8.99 (dd, 1H), 8.83 (d, 1H), 8.39 (d, 1H), 8.04 (m, 1H), 7.59 (m, 1H).	MS 249.98 (M ⁺ +H).
103		8.85 (d, 1H), 8.35 (d, 1H), 8.03 (d, 1H), 7.90-7.94 (m, 2H), 7.57 (dd, 1H), 7.45 (m, 1H), 7.15-7.17 (m, 1H), 4.88 (s, 2H).	MS 279.33 (M ⁺ +H).
104		8.94 (d, 1H), 8.77 (d, 1H), 8.45-8.53 (m, 2H), 8.38 (d, 1H), 8.30 (m, 1H), 8.22 (m, 1H), 8.05 (m, 1H), 7.79 (d, 1H), 7.60 (dd, 1H), 7.44 (m, 1H).	MS 317.21 (M ⁺ +H).
105		9.52 (m, 1H), 9.05-9.09 (m, 2H), 8.91-8.94 (m, 2H), 8.75-8.90 (m, 3H), 8.57 (m, 1H), 8.29 (m, 1H), 8.23 (m, 1H).	MS 318.39 (M ⁺ +H).
106		9.43 (s, 1H), 8.85-8.87 (m, 2H), 7.28 (s, 2H).	MS 292.11 (M ⁺ +H).
107		9.48 (s, 1H), 9.07 (m, 1H), 8.99 (s, 1H), 8.89-8.95 (m, 2H), 8.86 (dd, 1H), 8.28 (dd, 1H).	MS 302.97 (M ⁺ +H).
108		9.05 (d, 1H), 8.87 (d, 1H), 8.54-8.63 (m, 3H), 8.36 (d, 1H), 7.94 (dd, 1H), 7.61 (d, 1H), 7.48 (m, 1H), 7.43 (m, 1H), 7.23 (d, 1H).	MS 318.40 (M ⁺ +H).

EXAMPLE	Structure	¹ H NMR	MS (ESI)
109		9.05 (d, 1H), 8.85 (d, 1H), 8.62 (dd, 1H), 8.47 (m, 2H), 8.35 (d, 1H), 7.93 (dd, 1H), 7.65 (m, 1H), 7.46 (m, 1H), 7.25 (d, 1H).	MS 352.40 (M ⁺ +H).
110		9.03 (d, 1H), 8.84 (d, 1H), 8.57 (dd, 1H), 8.43 (d, 1H), 8.34 (d, 1H), 7.92 (dd, 1H), 7.46 (m, 2H), 7.19 (d, 1H), 7.15 (d, 1H), 2.61 (s, 3H).	MS 332.37 (M ⁺ +H).
111		9.02 (d, 1H), 8.85 (d, 1H), 8.58 (dd, 1H), 8.45 (d, 1H), 8.35 (d, 1H), 7.92 (dd, 1H), 7.46 (m, 1H), 7.25 (m, 1H), 7.15 (d, 1H), 2.46 (s, 3H).	MS 332.37 (M ⁺ +H).
112		9.12 (s, 1H), 8.94 (d, 1H), 8.71 (d, 1H), 8.47 (dd, 1H), 8.35 (d, 1H), 8.21 (d, 1H), 7.91 (dd, 1H), 7.36-7.46 (m, 3H), 6.87 (m, 1H), 2.35 (s, 3H).	MS 331.09 (M ⁺ +H).
113		9.46 (s, 1H), 8.95 (d, 2H), 8.90 (m, 1H), 8.78 (br, 1H), 8.39 (d, 1H), 8.16 (d, 1H), 8.09 (dd, 1H), 7.63 (dd, 1H), 6.94 (d, 2H).	MS 318.04 (M ⁺ +H).
114		8.86-8.87 (d, 1H), 8.62-8.63 (d, 1H), 8.41-8.42 (dd, 1H), 8.35-8.37 (d, 1H), 7.93-7.95 (m, 2H), 7.48-7.49 (t, 1H).	MS 328.7 (M ⁺ +2).
115		8.84-8.85 (d, 1H), 8.34-8.35 (d, 1H), 7.90-7.91 (t, 1H), 7.61-7.63 (d, 1H), 7.60 (s, 1H), 7.42-7.42 (m, 1H), 7.31-7.34 (t, 1H), 6.78-6.80 (d, 1H), 4.00 (s, 2H).	MS 239.1 (M ⁺ +1).
116		8.88-8.89 (d, 1H), 8.49 (s, 1H), 8.38-8.40 (d, 1H), 8.35-8.37 (d, 1H), 7.95-7.98 (m, 1H), 7.51-7.56 (m, 2H).	MS 268.1 (M ⁺ +1).

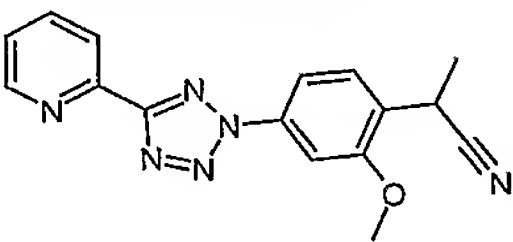
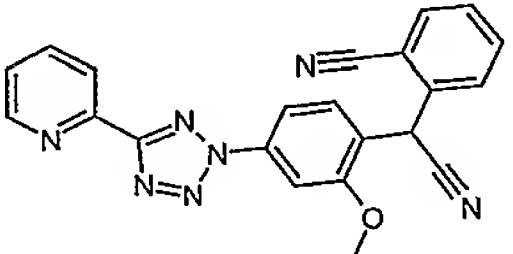
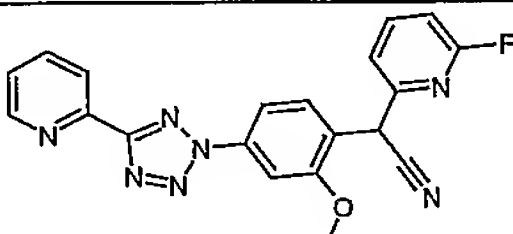
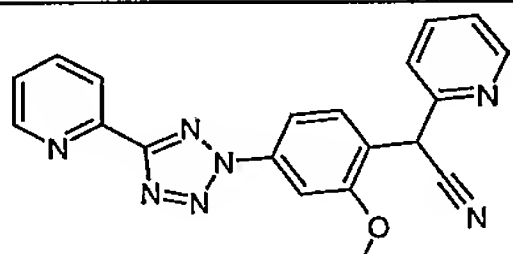
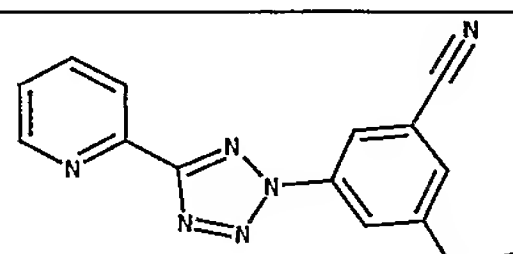
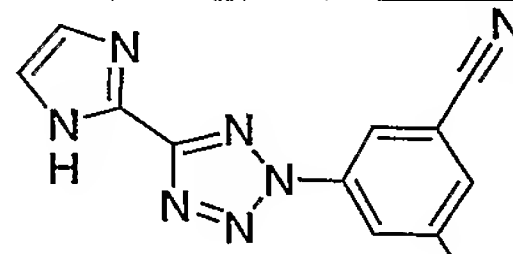
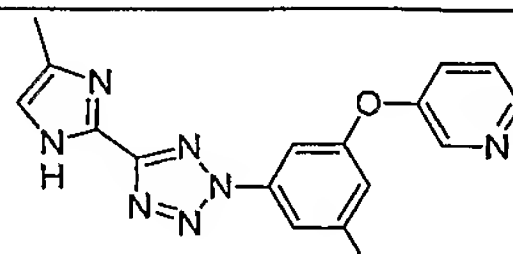
EXAMPLE	Structure	¹ H NMR	MS (ESI)
117		8.86-8.87(d, 1H), 8.70 (s, 1H), 8.54 (s, 1H), 8.37-8.39(d, 1H), 7.91-7.93(m, 2H), 7.48-7.49(m, 1H).	MS 370 (M ⁺ +2).
118		8.87-8.88(d, 1H), 8.84 (s, 2H), 8.36-8.37(d, 1H), 8.06 (s, 1H), 7.94-7.99(m, 1H), 7.49-7.52(m, 1H)	
119		9.39-9.40 (d, 1H), 8.97 (s, 1H), 8.88-8.89 (d, 1H), 8.65 (s, 1H), 8.39-8.41 (d, 1H), 7.92-7.98 (t, 1H), 7.47-7.53 (m, 1H).	MS 294.1 (M ⁺ +1)
120		8.80-8.81 (d, 1H), 8.25-8.27 (d, 1H), 8.05-8.09 (t, 1H), 7.70 (s, 1H), 7.61-7.63 (m, 2H), 7.08 (s, 1H), 6.28 (s, 2H).	MS 264.1 (M ⁺ +1)
121		8.87-8.88 (d, 1H), 8.60-8.61 (d, 1H), 8.53 (s, 1H), 8.37-8.39 (d, 1H), 7.94-7.98 (t, 1H), 7.78 (s, 1H), 7.50-7.52 (t, 1H).	MS 283.0 (M ⁺ +1)
122		8.59 (s, 1H), 8.50-8.52 (d, 1H), 8.43-8.46 (m, 1H), 8.22-8.23(d, 1H), 8.08-8.10 (dd, 1H), 7.64-7.68 (t, 1H), 7.47-7.61 (t, 1H).	MS 283.0 (M ⁺ +1)

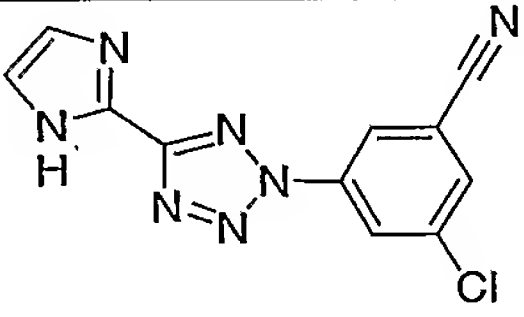
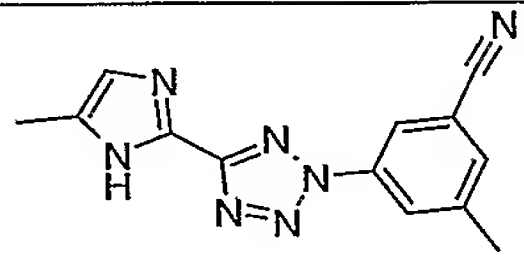
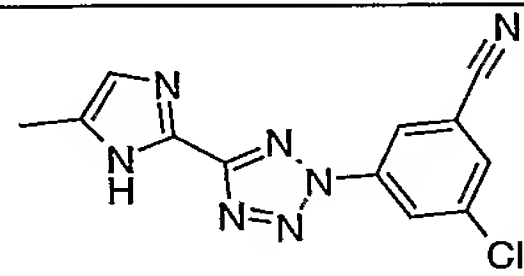
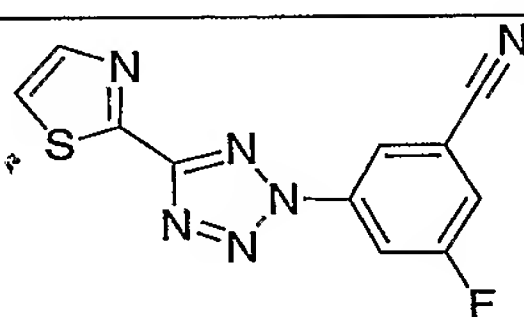
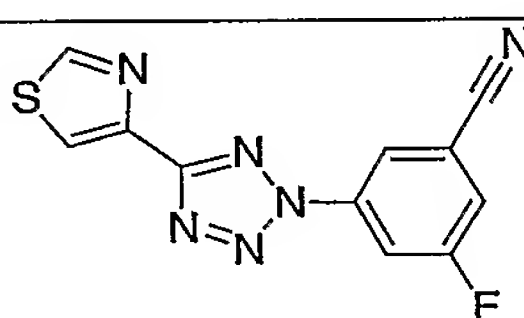
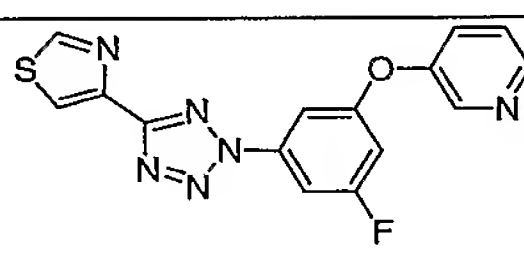
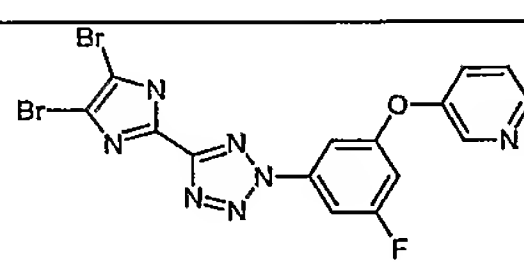
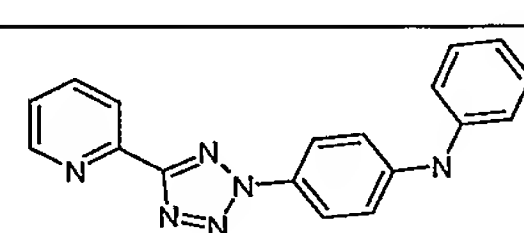
EXAMPLE	Structure	¹ H NMR	MS (ESI)
123		8.86-8.88 (d, 3H), 8.37-8.39 (d, 1H), 7.98-8.08 (s, 1H), 7.94-7.96 (t, 1H), 7.50-7.52 (m, 1H),	MS 274.0 (M ⁺ +H).
124		8.74-8.71 (m, 1H), 8.50-8.46 (m, 2H), 8.23 (d, 1H), 8.15-8.13 (m, 1H), 8.03 (td, 1H), 7.88-7.86 (m, 1H), 7.55-7.51 (m, 1H), 7.42-7.39 (m, 1H), 7.35 (dd, 1H), 7.19-7.15 (m, 1H)	MS 351.0 (M ⁺ +H).
125		8.73-8.70 (m, 1H), 8.37 (t, 1H), 8.30 (d, 1H), 8.26 (t, 1H), 8.16 (td, 1H), 7.72 (t, 1H), 7.66-7.63 (m, 1H).	MS 337.7 (M ⁺ +H).
126		8.73-8.70 (m, 1H), 8.58-8.56 (m, 1H), 8.55-8.53 (m, 1H), 8.31 (d, 1H), 8.19 (td, 1H), 7.95-7.94 (m, 1H), 7.68 (dd, 1H)	MS 282.9 (M ⁺ +H).
127		8.75 (d, 1H), 8.44 (s, 1H), 8.40 (s, 1H), 8.27 (d, 1H), 8.06 (td, 1H), 7.61 (s, 1H), 7.55 (dd, 1H), 2.51 (s, 3H)	MS 263.0 (M ⁺ +H).
128		8.82 (d, 1H), 8.29 (d, 1H), 8.18 (s, 1H), 8.06-8.11 (m, 2H), 7.63-7.68 (m, 2H), 7.56-7.57 (d, 1H), 4.68 (s, 2H).	MS 254.1 (M ⁺ +H).
129		8.82-8.83 (d, 1H), 8.29-8.30 (d, 1H), 8.24 (s, 1H), 8.18-8.19 (d, 1H), 8.07- 8.11 (dt, 1H), 7.74-7.78 (t, 1H), 7.62- 7.65 (m, 2H), 4.30 (s, 2H).	MS 263.1 (M ⁺ +H).

EXAMPLE	Structure	¹ H NMR	MS (ESI)
130		8.81-8.82 (d, 1H), 8.26-8.28 (d, 1H), 8.06-8.10 (dt, 1H), 7.61-7.64 (m, 1H), 7.38-7.41 (m, 1H), 7.34-7.36 (m, 2H), 6.80-6.82 (m, 1H), 2.79 (s, 3H).	MS 253.1 (M ⁺ +H).
131		9.23 (s, 1H), 8.77 (m, 1H), 8.59-8.62 (t, 1H), 8.43 (m, 1H), 8.36-8.38 (d, 1H), 8.09 (m, 1H), 7.80-7.83 (m, 2H), 7.72-7.74 (d, 1H), 7.68 (m, 1H), 5.71 (s, 2H).	MS 304.1 (M ⁺ +H).
132		8.97 (d, 1H), 8.87-8.88 (d, 1H), 8.70-8.73 (dt, 1H), 8.13-8.16 (t, 1H), 7.85-7.86 (t, 1H), 7.64-7.66 (m, 1H), 7.47-7.50 (t, 1H), 7.36-7.38 (m, 1H), 7.24-7.26 (m, 1H), 7.07-7.08 (m, 2H), 6.92-6.98 (m, 1H), 3.91 (s, 3H).	MS 345.1 (M ⁺ +H).
133		8.87-8.89 (m, 1H), 8.53 (s, 1H), 8.37-8.39 (d, 1H), 8.05-8.07 (td, 1H), 7.93-7.96 (m, 1H), 7.62-7.64 (m, 1H), 7.48-7.50 (m, 1H).	MS 368.4 (M ⁺ +H).
134		8.55 (s, 1H), 8.44-8.46 (d, 1H), 8.29-8.31 (d, 1H), 7.79 (m, 2H).	MS 256.4 (M ⁺ +H).
135		8.47 (s, 1H), 8.33-8.35 (m, 1H), 7.98-8.00 (m, 1H).	MS 414.1 (M ⁺ +H).
136		9.02 (s, 1H), 8.81-8.86 (m, 2H), 8.52-8.54 (d, 1H), 8.28-8.29 (d, 1H), 8.07-8.10 (t, 1H), 7.96-7.99 (t, 1H), 7.75 (s, 1H), 7.70-7.72 (d, 1H), 7.62-7.64 (t, 1H), 7.32-7.34 (d, 1H), 5.50 (s, 2H).	MS 349.3 (M ⁺ +H).

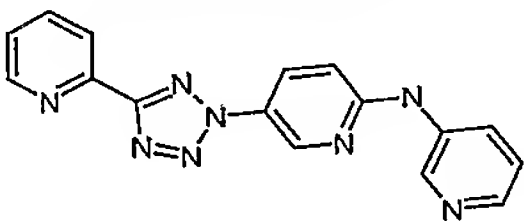
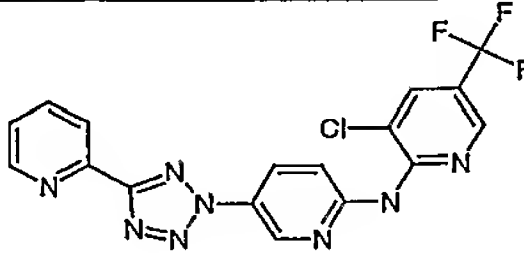
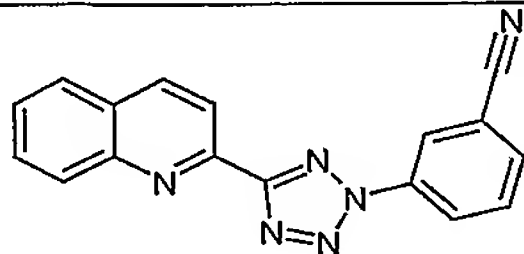
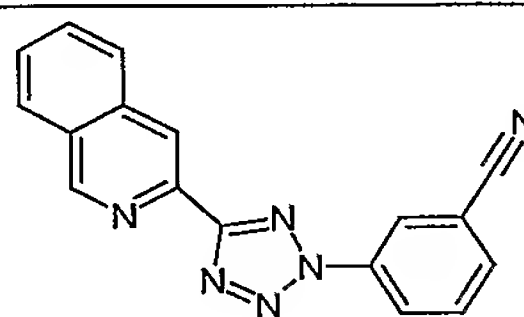
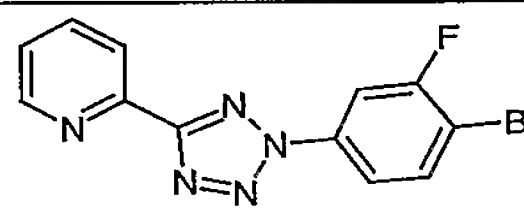
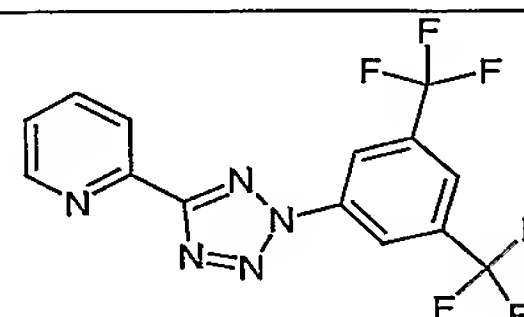
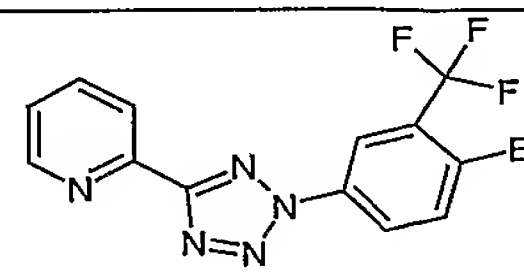
EXAMPLE	Structure	¹ H NMR	MS (ESI)
137		8.79-8.80 (m, 2H), 8.25-8.29 (m, 2H), 8.07-8.09 (t, 1H), 7.90-7.92 (d, 1H), 7.70-7.75 (m, 3H), 7.63 (t, 1H), 7.32-7.33 (d, 1H), 5.55 (s, 2H).	MS 373.3 (M ⁺ +H).
138		8.63 (m, 1H), 8.44 (m, 1H), 8.14 (m, 1H), 7.85 (m, 1H), 7.71-7.78 (m, 4H), 7.42-7.45 (m, 1H), 3.34 (s, 3H).	MS 337.5 (M ⁺ +H).
139		8.65-8.66 (m, 2H), 8.19 (m, 1H), 7.91 (s, 2H).	MS 272.7 (M ⁺ +H).
140		8.68-8.69 (d, 1H), 8.58-8.59 (d, 1H), 7.95-7.98 (dt, 1H), 7.87-7.89 (m, 2H), 7.74 (m, 1H), 7.66-7.68 (m, 2H), 7.41-7.44 (dt, 1H), 4.17 (s, 3H).	MS 338.4 (M ⁺ +H). ²
141		8.85-8.87 (d, 1H), 8.32-8.38 (m, 2H), 7.88-7.96 (m, 2H), 7.63-7.67 (m, 1H), 7.46-7.51 (m, 1H).	MS 283.1 (M ⁺ +H).
142		8.82-8.84 (d, 1H), 8.32-8.35 (d, 1H), 7.89-7.94 (m, 1H), 7.67-7.70 (m, 2H), 7.43-7.49 (m, 2H).	MS 292.1 (M ⁺).
143		8.81-8.83 (d, 1H), 8.31-8.34 (d, 1H), 7.87-7.92 (t, 1H), 7.69-7.70 (d, 1H), 7.41-7.45 (m, 2H), 7.07-7.10 (m, 1H), 3.88 (s, 3H).	MS 288.1 (M ⁺ +H).
144		ND	MS 249.2 (M ⁺ +H).

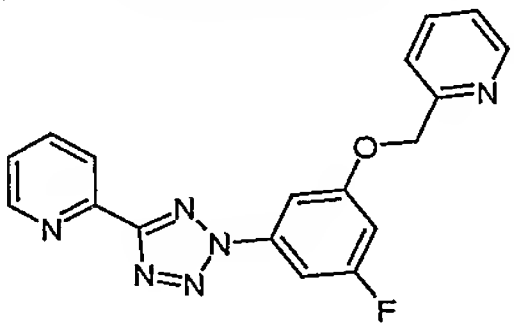
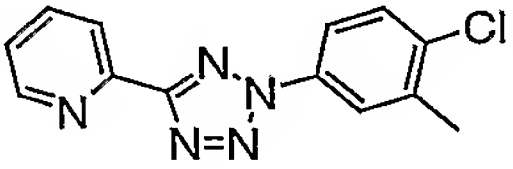
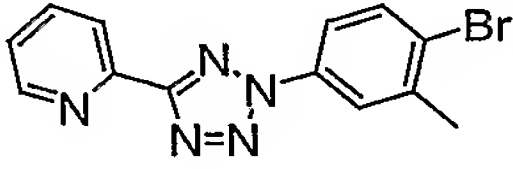
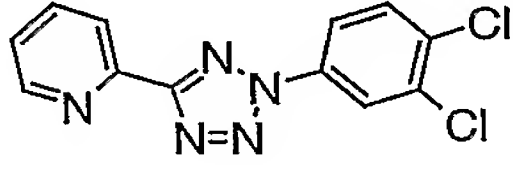
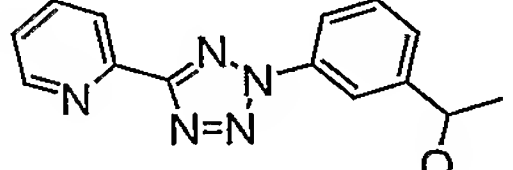
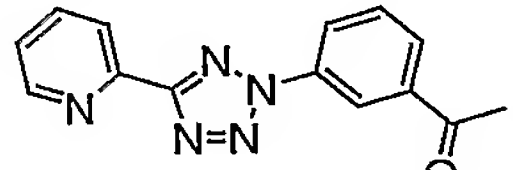
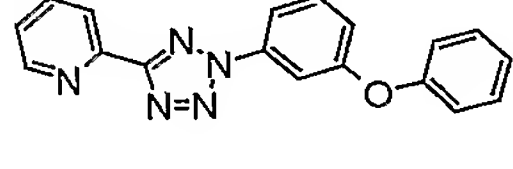
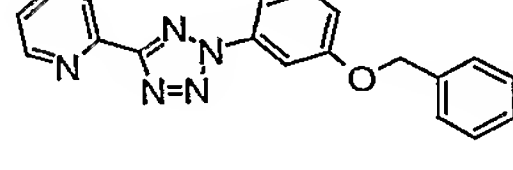
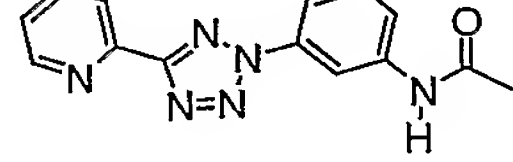
EXAMPLE	Structure	¹ H NMR	MS (ESI)
145		ND	MS 224.2 (M ⁺ +H).
146		8.85-8.86 (d, 1H), 8.36-8.38 (d, 1H), 7.91-7.94 (m, 2H), 7.81 (s, 1H), 7.57-7.59 (d, 1H), 7.45-7.48 (m, 1H), 4.02 (s, 3H), 3.78 (s, 1H).	MS 293.3 (M ⁺ +H).
147		8.83-8.84 (d, 1H), 8.34-8.36 (d, 1H), 7.87-7.91 (t, 1H), 7.75-7.77 (m, 1H), 7.69 (s, 1H), 7.43-7.44 (m, 1H), 7.28-7.29 (d, 1H), 3.96 (s, 3H), 2.28 (s, 3H).	MS 268.3 (M ⁺ +H).
148		8.85-8.86 (d, 1H), 8.36-8.38 (d, 1H), 7.89-7.92 (m, 2 H), 7.87 (s, 1H), 7.53-7.55 (d, 1H), 7.46-7.48 (m, 1H), 4.09 (s, 3H), 1.83 (s, 6H).	MS 321.3 (M ⁺ +H).
149		8.84-8.85 (m, 2H), 8.36-8.38 (d, 1H), 7.99-8.03 (dd, 2H), 7.88-7.92 (t, 2H), 7.81 (s, 1H), 7.39-7.49 (m, 2H), 6.14 (s, 1H), 4.01 (s, 3H).	MS 394.9 (M ⁺ +H).
150		9.11 (br, 1H), 8.72 (br, 1H), 8.49 (br, 1H), 8.18 (s, 1H), 8.05-8.06, (d, 1H), 7.97 (br, 1H), 7.76-7.78 (d, 1H), 4.61-4.63 (t, 1H), 4.15 (s, 3H), 3.00-3.12 (dq, 2H).	MS 332.0 (M ⁺ +H).
151		9.07-9.08 (d, 1H), 8.70-8.71 (d, 1H), 8.45-8.48 (t, 1H), 8.33 (s, 1H), 8.10-8.12 (d, 1H), 7.93-7.95 (t, 1H), 7.90-7.92 (d, 1H), 4.22 (s, 3H), 3.46-3.55 (q, 4H).	MS 371.0 (M ⁺ +H).

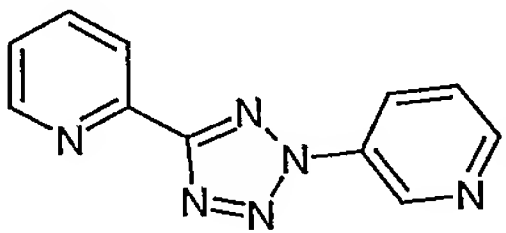
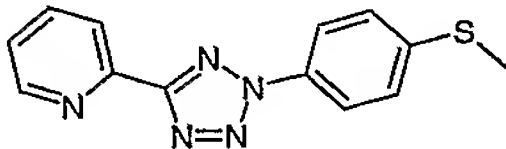
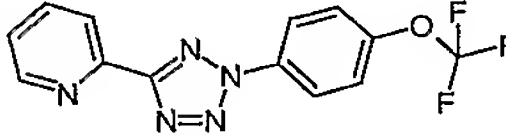
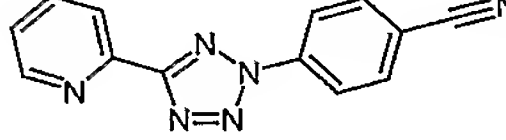
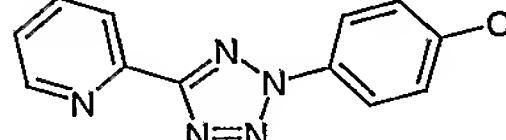
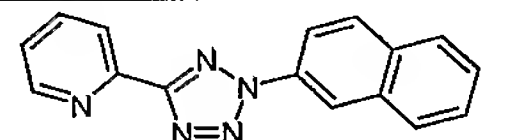
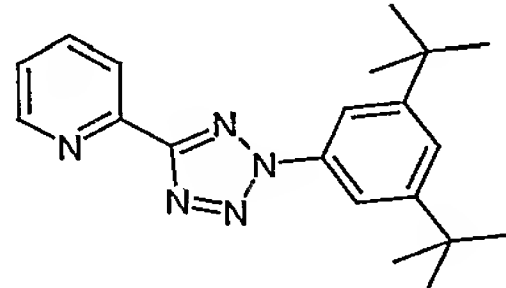
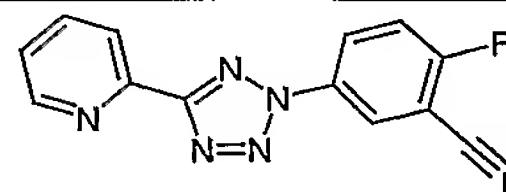
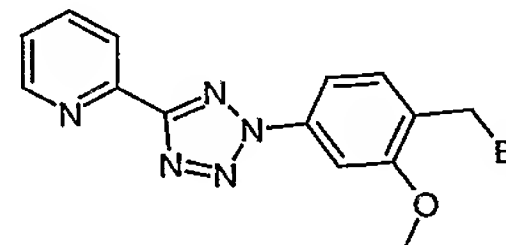
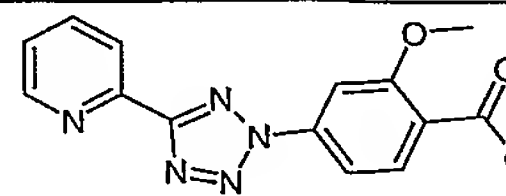
EXAMPLE	Structure	¹ H NMR	MS (ESI)
152		8.73 (s, 1H), 8.34-8.36 (d, 1H), 8.05-8.34 (t, 1H), 7.87-7.89 (d, 1H), 7.87 (s, 1H), 7.66-7.67 (d, 1H), 7.59-7.61 (m, 1H).	MS 307.0 (M ⁺ +H).
153		8.86 (s, 1H), 8.36-8.38 (d, 1H), 7.96-7.97 (d, 1H), 7.91-7.94 (t, 1H), 7.83-7.84 (t, 1H), 7.72-7.74 (d, 1H), 7.67-7.68 (t, 2H), 7.61-7.63 (d, 1H), 7.46-7.50 (m, 2H), 5.88 (s, 1H), 4.00 (s, 3H).	MS 394.0 (M ⁺ +H).
154		8.85-8.86 (d, 1H), 8.36-8.38 (d, 1H), 7.96-7.98 (dd, 1H), 7.91-7.94 (t, 1H), 7.82-7.86 (m, 2H), 7.65-7.67 (d, 1H), 7.44-7.47 (m, 1H), 7.32-7.34 (d, 1H), 6.92-6.94 (d, 1H) 5.64 (s, 1H), 3.99 (s, 3H)	MS 388.0 (M ⁺ +H).
155		8.85-8.86 (d, 1H), 8.61-8.62 (d, 1H), 8.36-8.38 (d, 1H), 7.91-7.96 (m, 2H), 7.84 (s, 1H), 7.72-7.75 (t, 1H), 7.64-7.66 (d, 1H), 7.45-7.47 (m, 1H), 7.40-7.42 (d, 1H), 5.73 (s, 1H), 4.00 (s, 3H)	MS 370.0 (M ⁺ +H).
156		8.65-8.66 (d, 1H), 8.22-8.60 (d, 1H), 8.0 (s, 1H), 7.90-7.97 (m, 2H), 7.49-7.52 (m, 1H), 7.43 (s, 1H)	MS 278.5 (M ⁺ +H).
157		8.42 (s, 1H), 8.34 (s, 1H), 8.02 (s, 1H), 7.71 (s, 2H), 3.17 (s, 3H).	MS 252.3 (M ⁺ +H).
158		8.80-8.81 (d, 1H), 8.66-8.68 (d, 1H), 8.08-8.10 (m, 1H), 7.89-7.92 (m, 1H), 7.82-7.85 (m, 1H), 7.72 (s, 1H), 7.67 (m, 1H), 7.50-7.53 (m, 1H), 2.40 (s,	MS 338.4 (M ⁺ +H).

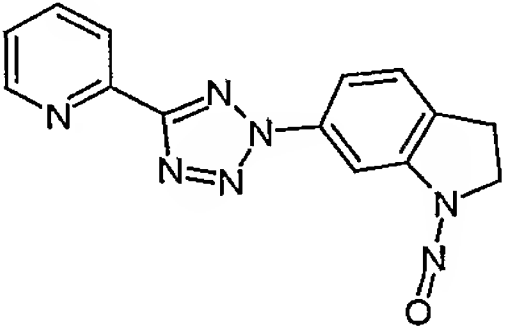
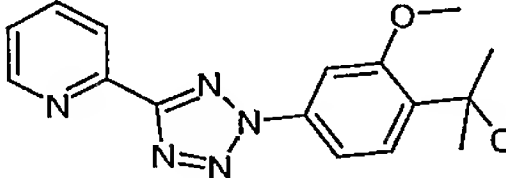
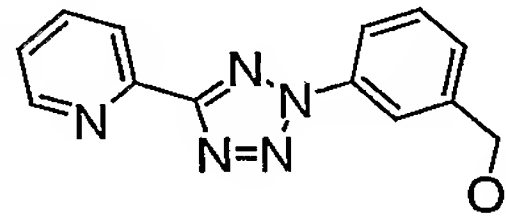
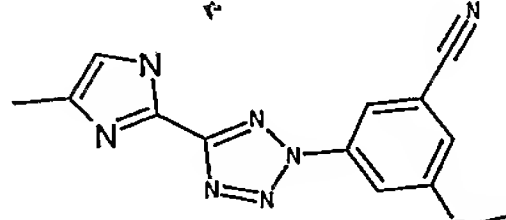
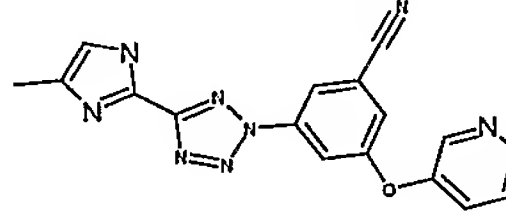
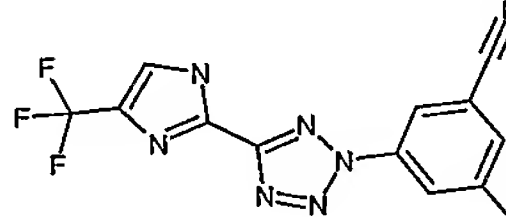
EXAMPLE	Structure	¹ H NMR	MS (ESI)
		3H).	
159		8.65-8.66 (m, 2H), 8.18 (s, 1H), 7.91 (m, 2H).	MS 272.3 (M ⁺ +H).
160		8.42 (s, 1H), 8.33 (s, 1H), 8.05 (s, 1H), 7.62 (s, 1H), 2.54 (s, 3H), 2.39 (s, 3H).	MS 266.27 (M ⁺ +H).
161		8.64 (s, 1H), 8.53 (s, 1H), 8.44 (s, 1H), 7.63 (s, 1H), 2.39 (s, 3H).	MS 286.18 (M ⁺ +H).
162		8.43 (s, 1H), 8.31-8.28 (m, 1H), 8.13 (d, 1H), 7.65 (d, 1H), 7.56-7.51 (m, 1H).	MS 273.26 (M ⁺ +H).
163		9.04 (d, 1H), 8.43 (s, 1H), 8.35 (d, 1H), 8.31-8.29 (d, 1H), 7.54-7.51 (m, 1H).	MS 273.21 (M ⁺ +H).
164		9.01 (d, 1H), 8.52-8.51 (m, 2H), 8.30 (d, 1H), 7.82-7.91 (m, 1H), 7.73 (s, 1H), 7.47-7.44 (m, 1H), 7.40-7.37 (m, 1H), 6.87-6.84 (m, 1H).	MS 341.21 (M ⁺ +H).
165		8.58 (d, 1H), 8.52 – 8.53 (m, 1H), 7.82 – 7.85 (m, 1H), 7.21 – 7.75 (m, 1H), 7.55 (dd, 1H), 7.54 (m, 1H), 7.34 – 7.37 (m, 1H).	MS 482.1 (M ⁺ +H).
166		8.92 (m, 1H), 8.74-8.73 (d, 1H), 8.66-8.63 (t, 1H), 8.07-8.07 (d, 3H), 7.34-7.31 (m, 2H), 7.26-7.21 (m, 4H), 7.02-	MS 315.00 (M ⁺).

EXAMPLE	Structure	¹ H NMR	MS (ESI)
		7.00 (m, 1H).	
167		8.84-8.83 (d, 1H), 8.34-8.332 (d, 1H), 8.02-8.00 (m, 1H), 7.94-7.93 (m, 1H), 7.93-7.89 (m, 1H), 7.53 (t, 1H), 7.45-7.43 (m, 1H), 7.29 (d, 1H), 7.17-7.15 (dd, 1H), 6.74-6.72 (dd, 1H), 6.68-6.63 (m, 2H), 3.80 (s, 3H).	MS 346.19 (M ⁺ +H).
168		8.53-8.52 (m 2H), 7.88-7.87 (d, 1H), 7.91-7.72 (m, 3H), 7.48-7.45 (ddd, 1H), 7.40-7.38 (dd, 1H), 6.91-6.90 (d, 1H), 6.85-6.83 (m, 1H), 4.09 (s, 3H).	MS 365.32 (M ⁺ +H).
169		9.56 (d, 1H), 8.85 (d, 1H), 8.79 (m, 1H), 8.57 (m, 1H), 8.36 (d, 1H), 7.93 (dd, 1H), 7.57 (m, 1H), 7.48 (m, 1H).	MS 225.1 (M ⁺ +H).
170		9.35 (d, 1H), 8.86 (d, 1H), 8.55 (dd, 1H), 8.36 (d, 1H), 7.94 (dd, 1H), 7.58 (d, 1H), 7.49 (m, 1H).	MS [*] 259.32 (M ⁺ +H).
171		9.46 (d, 1H), 8.77 (d, 1H), 8.61 (d, 1H), 7.69 (dd, 1H), 7.30 (s, 2H).	MS 214.12 (M ⁺ +H).
172		8.36 (d, 1H), 8.32 (d, 1H), 8.02 (m, 2H), 7.89 (dd, 1H), 7.70 (d, 1H), 7.51 (m, 1H), 7.39 (m, 1H).	MS 224.99 (M ⁺ +H).
173		8.89 (s, 1H), 8.72 (d, 1H), 8.22 (m, 1H), 8.10 (m, 1H), 7.79 (m, 1H), 7.29 (m, 1H), 6.48 (m, 1H), 3.05 (s, 6H).	MS 268.39 (M ⁺ +H).
174		9.29 (d, 1H), 9.15 (d, 1H), 8.77-8.84 (m, 2H), 8.69 (m, 1H), 8.36 (d, 1H), 8.18-8.26 (m, 2H), 7.49 (m, 2H), 7.24 (dd, 1H).	MS 316.94 (M ⁺ +H).

EXAMPLE	Structure	¹ H NMR	MS (ESI)
175		ND	MS 316.94 (M ⁺ +H).
176		9.22 (m, 1H), 8.86 (d, 1H), 8.78 (d, 1H), 8.57 (m, 1H), 8.53 (s, 1H), 8.35 (d, 1H), 8.31 (s, 1H), 7.90-7.95 (m, 2H), 7.46 (m, 1H).	MS 418.86 (M ⁺ +H).
177		9.34-9.37(d, 1H), 8.87-8.90(d, 1H), 8.74(s, 1H), 8.66-8.68(d, 1H), 8.57- 8.60(d, 1H), 8.41-8.44(d, 1H), 8.26- 8.29(t, 1H), 7.95-8.08(m, 3H).	MS 299.1 (M ⁺ +1)
178		9.55(s, 1H), 8.84(s, 1H), 8.66(s, 1H), 8.53-8.55(d, 1H), 8.23-8.29(m, 2H), 8.11-8.14(d, 1H), 7.81-7.96(m, 3H).	MS 299.1 (M ⁺ +1)
179		8.84-8.88(d, 1H), 8.32-8.36(d, 1H), 8.12-8.14(d, 1H), 8.03-8.05(d, 1H), 7.93-7.96(t, 1H), 7.77-7.82(t, 1H), 7.46-7.49(m, 1H).	MS 321.1 (M ⁺ +1)
180		9.02-9.03(d, 1H), 8.86-8.87(m, 2H), 8.74-8.76(t, 1H), 8.34 (s, 1H), 8.17-8.19(t, 1H).	MS 360.1 (M ⁺ +1)
181		8.75-8.76(d, 1H), 8.60-8.61(d, 1H), 8.35-8.41(m, 2H), 8.12-8.14(d, 1H), 8.06-8.09(t, 1H), 7.60-7.62(m, 1H).	MS 371.1 (M ⁺ +1)

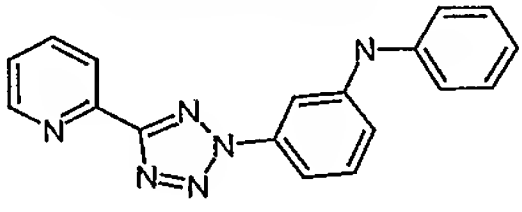
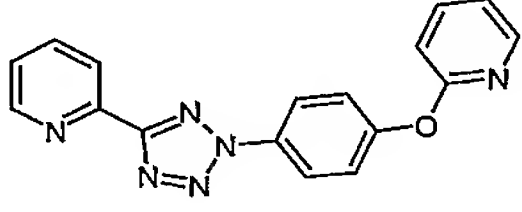
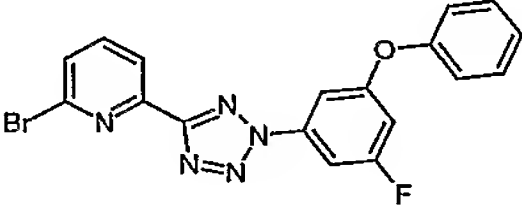
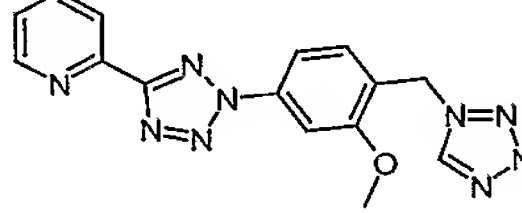
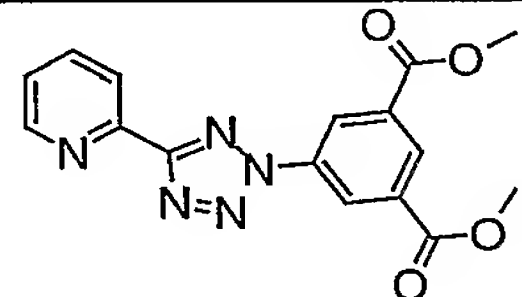
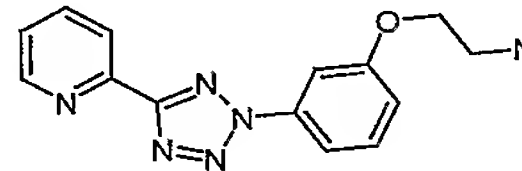
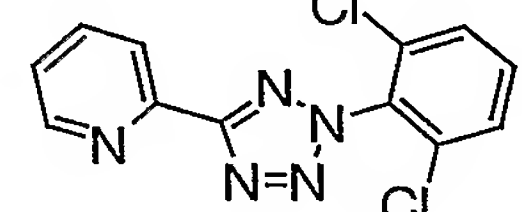
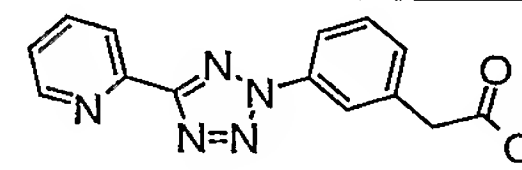
EXAMPLE	Structure	¹ H NMR	MS (ESI)
182		8.81 (d, 1H), 8.27-8.28 (d, 1H), 8.06-8.09 (t, 1H), 7.94-7.95 (d, 1H), 7.78-7.80 (m, 2H), 7.67-7.70 (m, 2H), 7.60-7.62 (m, 2H), 7.30-7.32 (d, 1H).	MS 349.3 (M ⁺ +H).
183		8.84-8.86 (d, 1H), 8.34-8.36 (d, 1H), 8.19-8.20 (d, 1H), 8.04-8.07 (m, 1H), 7.88-7.94 (m, 1H), 7.52-7.55 (d, 1H), 7.43-7.47 (m, 1H), 2.50 (s, 3H).	MS 272.2 (M ⁺ +H).
184		ND	MS 316.0 (M ⁺).
185		ND	MS 292.0 (M ⁺).
186		8.83-8.85 (d, 1H), 8.34-8.37 (d, 1H), 8.29 (s, 1H), 8.14-8.18 (m, 1H), 7.89-7.94 (m, 1H), 7.53-7.55 (m, 2H), 7.43-7.47 (m, 1H), 5.02-5.09 (q, 1H), 2.34 (s, 1H), 1.57-1.59 (d, 3H).	MS 268.0 (M ⁺ +H).
187		8.85-8.90 (m, 2H), 8.50-8.54 (m, 1H), 8.39-8.42 (m, 1H), 8.13-8.16 (m, 1H), 7.95-7.97 (m, 1H), 7.71-7.75 (m, 1H), 7.49-7.51 (m, 1H), 2.75 (s, 3H).	MS 266.1 (M ⁺ +H).
188		8.84-8.86 (m, 1H), 8.34-8.37 (d, 1H), 8.00-8.03 (m, 1H), 7.92-7.95 (m, 2H), 7.39-7.57 (m, 4H), 7.10-7.18 (m, 4H).	MS 316.1 (M ⁺ +H).
189		ND	MS 330.1 (M ⁺ +H).
190		ND	MS 281.0 (M ⁺ +H).

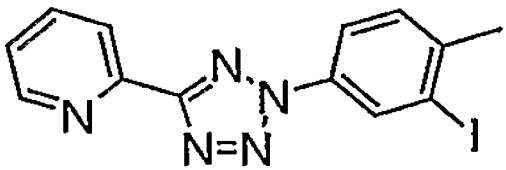
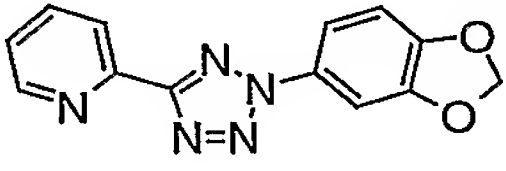
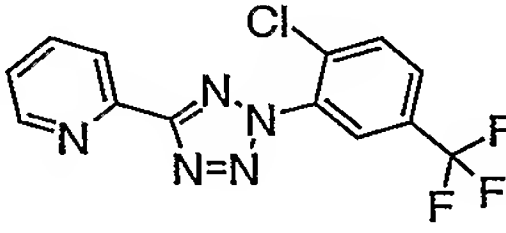
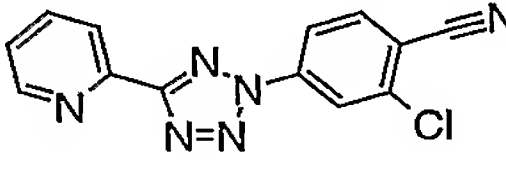
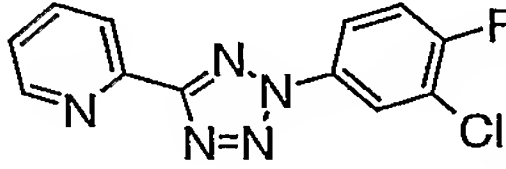
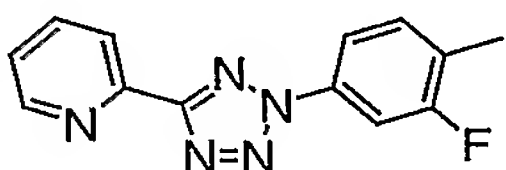
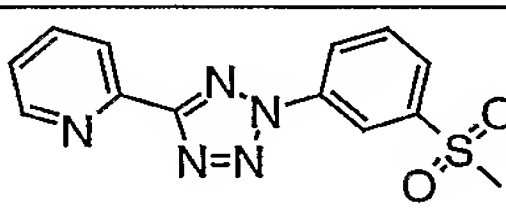
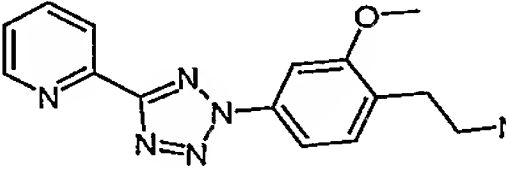
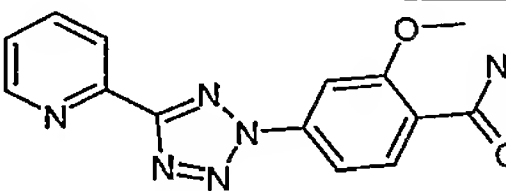
EXAMPLE	Structure	¹ H NMR	MS (ESI)
191		ND	MS 225.1 (M ⁺ +H).
192		ND	MS 270.2 (M ⁺ +H).
193		ND	MS 308.2 (M ⁺ +H).
194		ND	MS 249.2 (M ⁺ +H).
195		ND	MS 258.5 (M ⁺ +H).
196		ND	MS 274.2 (M ⁺ +H).
197		ND	MS 335.3 (M ⁺ +H).
198		ND	MS 267.2 (M ⁺ +H).
199		8.85-8.86 (m, 1H), 8.36-8.37 (d, 1H), 7.91-7.93 (t, 1H), 7.86-7.87 (d, 1H), 7.80 (s, 1H), 7.52-7.54 (d, 1H), 7.44- 7.46 (m, 1H), 4.59 (s, 2H), 4.04 (s, 3H).	MS 346.0 (M ⁺ +H).
200		8.83 (s, 1H), 8.30-8.32 (d, 1H), 8.07- 8.10 (t, 1H), 7.96-7.98 (d, 1H), 7.83- 7.86 (m, 2H), 7.62-7.63 (m, 1H), 4.02	MS 298.1 (M ⁺ +H).

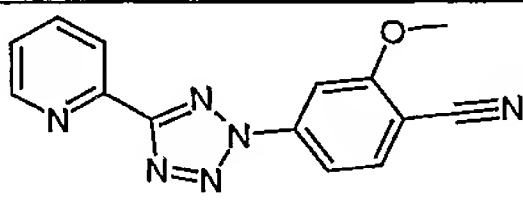
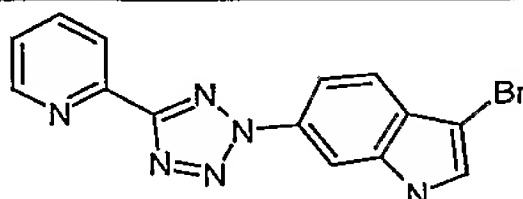
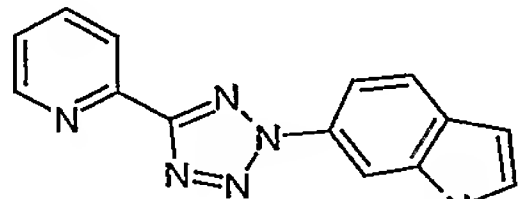
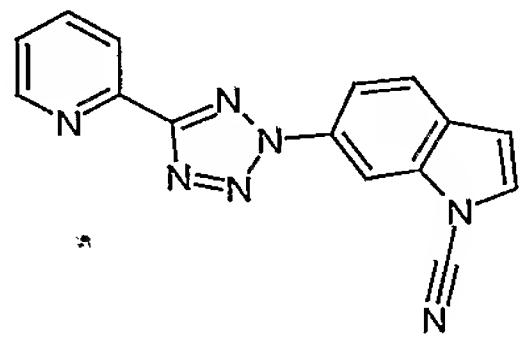
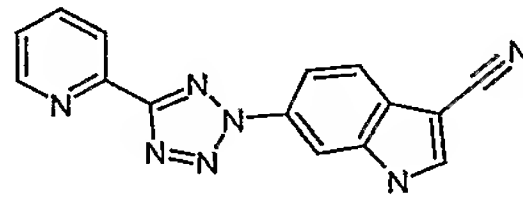
EXAMPLE	Structure	¹ H NMR	MS (ESI)
		(s, 3H).	
201		8.83-8.84 (d, 1H), 8.70 (s, 1H), 8.33-8.34 (d, 1H), 8.12-8.14 (dd, 1H), 7.90-7.93 (t, 1H), 7.51-7.52 (d, 1H), 7.45-7.47 (dd, 1H), 4.21-4.24 (t, 2H), 3.30-3.33 (t, 2H).	MS 294.2 (M ⁺ +H).
202		8.84-8.85 (dd, 1H), 8.35-8.36 (d, 1H), 7.89-7.91 (t, 1H), 7.84-7.86 (dd, 1H), 7.82 (s, 1H), 7.57-7.59 (d, 1H), 7.43-7.45 (m, 1H), 4.05 (s, 3H), 1.67 (s, 6H).	MS 312.1 (M ⁺ +H).
203		8.86-8.88 (d, 1H), 8.37-8.40 (d, 1H), 8.33 (s, 1H), 8.21-8.23 (m, 1H), 7.91-7.96 (m, 1H), 7.54-7.61 (m, 2H), 7.45-7.48 (m, 1H), 4.86-4.88 (d, 2H), 1.92-1.96 (t, 1H).	MS 254.1 (M ⁺ +H).
204		8.18(s, 1H), 8.07 (s, 1H), 7.54(s, 1H), 7.19(s, 1H), 4.01(s, 3H), 2.36(s, 3H).	*MS 282.4 (M ⁺ +H).
205		8.51-8.50(d, 1H), 8.48-8.47(d, 1H), 8.39(s, 1H), 8.12-8.11(d, 1H), 7.71-7.69(d, 1H), 7.63(d,1H), 7.56-7.54(m,1H), 7.19(s, 1H), 2.39(s, 3H)	MS 346.1 (M ⁺ +H).
206		8.47 (s, 1H), 8.36 (m, 1H), 7.67 (m, 2H).	MS 324.17 (M ⁺ +H).

Examples 207-236 have mGluR5 inhibitory activity greater than 10 μ M in the calcium flux assay or inhibition <50% at 100 μ M concentration in the PI assay.

EXAMPLE	Structure	¹ H NMR	MS (ESI)
207		8.71 (s, 1H), 8.52 – 8.54 (m, 1H), 8.13 – 8.15 (m, 1H), 7.92 (dd, 1H), 7.68 (d, 1H), 7.07 (d, 1H), 4.31 (s, 3H)	MS 252.1 (M ⁺ +H).
208		8.96 (s, 1H), 8.29 (s, 1H), 8.25 (m, 1H), 8.16 – 8.17 (m, 1H), 7.81 – 7.83 (m, 1H), 7.75 – 7.78 (dd, 1H).	MS 238.1 (M ⁺ +H).
209		8.63 – 8.64 (m, 1H), 8.49 – 8.51 (m, 1H), 8.38 – 8.39 (m, 1H), 8.12 – 8.14 (m, 1H), 7.93 (dd, 1H), 7.78 – 7.80 (m, 1H), 7.66 (dd, 1H), 2.51 (m, 3H).	MS 279.1 (M ⁺ +H).
210		8.68 – 8.69 (m, 1H), 8.54 – 8.55 (m, 1H), 8.17 – 8.18 (m, 1H), 7.97 (dd, 1H), 7.72 – 7.74 (m, 2H), 7.35 – 7.37 (m, 2H).	MS 288.2 (M ⁺ +H).
211		8.71 – 8.72 (m, 1H), 8.67 (m, 1H), 8.59 – 8.61 (m, 1H), 8.06 – 8.10 (m, 1H), 7.77 – 7.81 (m, 1H).	MS 365.0 (M ⁺ +H).
212		8.71 (m, 1H), 8.58 – 8.60 (m, 1H), 8.15 (d, 1H), 8.00 – 8.02 (m, 1H), 7.90 – 7.94 (m, 2H), 7.28 (m, 1H), 7.66 – 7.69 (m, 3H), 7.40 – 7.43 (m, 2H), 6.87 (d, 2H).	MS 374.4 (M ⁺ +H).
213		9.01 (d, 1H), 8.84 (d, 1H), 8.76 (ddd, 1H), 8.20 (ddd, 1H), 7.04 (s, 1H), 2.63 (s, 3H).	MS 229.1 (M ⁺ +H).
214		8.79 (br, 1H), 8.41 (d, 3H), 8.14-8.11 (m, 2H), 8.01 (d, 1H), 7.72 (d, 2H), 7.65 (br, 1H), 7.34 (d, 1H), 7.16 (t, 1H).	MS 315.90 (M ⁺ +H).

EXAMPLE	Structure	¹ H NMR	MS (ESI)
215		8.92 (br 1H), 8.71-8.69 (m, 1H), 8.64-8.62 (m, 1H), 8.07 (br, 1H), 7.85 (br, 1H), 7.59-7.57 (m, 1H), 7.44-7.41 (m, 1H), 7.31-7.28 (m, 2H), 7.21-7.17 (m, 3H), 6.98-6.95 (m, 1H).	MS 315.22 (M ⁺ +H).
216		9.03 (br, 1H), 8.90-8.88 (m, 1H), 8.84-8.80 (m, 1H), 8.45-8.80 (m, 3H), 8.27-8.19 (m, 2H), 7.64-7.62 (d, 2H), 7.50-7.48 (m, 1H), 7.29-7.28 (m, 1H).	MS 316.49 (M ⁺ +H).
217		8.524-8.515 (m, 2H), 8.27-8.26 (dd, 1H), 7.80-7.73 (m, 2H), 7.729-7.727 (m, 1H), 7.66-7.64 (dd, 1H), 7.47-7.46 (m, 1H), 7.41-7.39 (m, 1H), 6.88-6.85 (m, 1H).	MS 413.30 (M ⁺ +H).
218		9.28 (s, 1H), 8.97 (m, 1H), 8.82-8.80 (m, 1H), 8.72-8.69 (t, 1H), 8.14-8.12 (t, 1H), 7.94-7.92 (m, 2H), 7.68-7.66 (d, 1H), 5.78 (s, 2H), 4.07 (s, 3H)	MS 336.3 (M ⁺ +H)
219		9.09-9.10 (s, 2H), 8.86-8.87 (d, 1H), 8.80-8.81 (d, 1H), 8.37-8.38 (d, 1H), 7.93-7.94 (t, 1H), 7.32-7.49 (m, 1H), 3.87-4.17 (s, 9H),	MS 340.0 (M ⁺ +H).
220		8.97-8.98 (d, 1H), 8.75-8.78 (d, 1H), 8.62-8.66 (dt, 1H), 8.07-8.10 (t, 1H), 7.95-7.97 (m, 2H), 7.67-7.70 (t, 1H), 7.33-7.36 (m, 1H), 4.42-4.44 (m, 2H), 3.48-3.50 (t, 2H).	MS 283.0 (M ⁺ +H).
221		8.83-8.85 (d, 1H), 8.35-8.38 (d, 1H), 7.90-7.96 (m, 1H), 7.51-7.60 (m, 3H), 7.44-7.48 (m, 1H).	MS 292.1 (M ⁺).
222		8.80-8.82 (d, 1H), 8.27-8.30 (d, 1H), 8.05-8.14 (m, 3H), 7.60-7.65 (m, 2H),	MS 282.0 (M ⁺ +H).

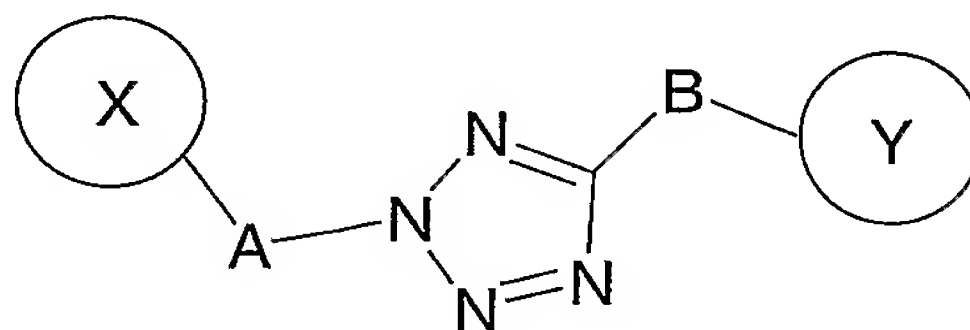
EXAMPLE	Structure	¹ H NMR	MS (ESI)
		7.52-7.54 (d, 1H), 3.82 (s, 2H).	
223		8.84-8.86 (d, 1H), 8.74-8.75 (d, 1H), 8.34-8.37 (d, 1H), 8.15-8.19 (m, 1H), 7.89-7.94 (m, 1H), 7.41-7.48 (m, 2H), 2.52 (s, 3H).	MS 364.0 (M ⁺ +H).
224		8.83-8.85 (d, 1H), 8.32-8.35 (d, 1H), 7.88-7.93 (m, 1H), 7.74-7.82 (m, 2H), 7.42-7.46 (m, 1H), 6.94-6.97 (d, 1H), 6.11 (s, 2H).	MS 268.2 (M ⁺ +H).
225		8.86-8.88 (d, 1H), 8.66-8.67 (d, 1H), 8.37-8.47 (m, 2H), 7.93-7.98 (m, 1H), 7.76-7.78 (d, 1H), 7.42-7.47 (m, 1H).	MS 326.1 (M ⁺ +H).
226		ND	MS 283.1 (M ⁺ +H).
227		ND	MS 276.0 (M ⁺ +H).
228		ND	MS 256.0 (M ⁺).
229		8.86-8.90 (m, 2H), 8.60-8.63 (m, 1H), 8.37-8.40 (m, 1H), 8.11-8.14 (m, 1H), 7.93-7.98 (m, 1H), 7.82-7.87 (t, 1H), 7.47-7.51 (m, 1H), 3.18 (s, 3H).	MS 302.0 (M ⁺ +H).
230		8.84-8.85 (m, 1H), 8.35-8.36 (d, 1H), 7.89-7.91 (t, 1H), 7.79-7.81 (dd, 1H), 7.73-7.74 (d, 1H), 7.43-7.45 (m, 1H), 7.32-7.33 (d, 1H), 3.96 (s, 3H), 2.98- 3.01 (t, 2H), 2.84-2.87 (t, 2H).	MS 297.2 (M ⁺ +H).
231		8.81-8.83 (m, 1H), 8.29-8.30 (d, 1H), 8.07-8.09 (t, 1H), 8.04-8.06 (d, 1H),	MS 297.0 (M ⁺ +H).

EXAMPLE	Structure	¹ H NMR	MS (ESI)
		4.06 (s, 3H).	
232		8.86-8.87 (m, 1H), 8.37-8.38 (d, 1H), 7.98-7.80 (dd, 1H), 7.93-7.95 (t, 1H), 7.92 (s, 1H), 7.78-7.79 (d, 1H), 7.49- 7.51 (m, 1H), 4.10 (s, 3H).	MS 279.0 (M ⁺ +H).
233		ND	MS 341.0 (M ⁺ +H).
234		11.74 (s, 1H), 8.82-8.83 (dd, 1H), 8.28- 8.29 (d, 1H), 8.22 (s, 1H), 8.06-8.09 (dt, 1H), 7.83-7.85 (m, 1H), 7.84 (s, 1H), 7.60-7.63 (m, 2H), 6.62 (s, 1H).	MS 263.1 (M ⁺ +H).
235		8.86-8.87 (d, 1H), 8.57 (s, 1H), 8.37- 8.39 (d, 1H), 8.30-8.32 (dd, 1H), 7.92- 7.94 (t, 1H), 7.82-7.84 (d, 1H), 7.46- 7.48 (dd, 1H), 7.41 (s, 1H), 6.64 (s, 1H).	MS 288.0 (M ⁺ +H).
236		12.67 (s, 1H), 8.81-8.83 (m, 1H), 8.50 (s, 1H), 8.35-8.36 (d, 1H), 8.28-8.30 (dd, 1H), 8.08-8.10 (m, 2H), 7.94-7.96 (d, 1H), 7.62-7.64 (m, 1H)	MS 288.0 (M ⁺ +H).

Other variations or modifications, which will be obvious to those skilled in the art, are within the scope and teachings of this invention. This invention is not to be limited except as set forth in the following claims.

WHAT IS CLAIMED IS:

1. A compound represented by Formula (I):



(I)

or a pharmaceutically acceptable salt thereof, wherein

X and Y each independently is aryl or heteroaryl wherein at least one of X and Y is a heteroaryl with N adjacent to the position of attachment to A or B respectively;

X is optionally substituted with 1-7 independent halogen, $-\text{CN}$, NO_2 , $-\text{C}_{1-6}\text{alkyl}$, $-\text{C}_{1-6}\text{alkenyl}$, $-\text{C}_{1-6}\text{alkynyl}$, $-\text{OR}^1$, $-\text{NR}^1\text{R}^2$, $-\text{C}(=\text{NR}^1)\text{NR}^2\text{R}^3$, $-\text{N}(=\text{NR}^1)\text{NR}^2\text{R}^3$, $-\text{NR}^1\text{COR}^2$, $-\text{NR}^1\text{CO}_2\text{R}^2$, $-\text{NR}^1\text{SO}_2\text{R}^4$, $-\text{NR}^1\text{CONR}^2\text{R}^3$, $-\text{SR}^4$, $-\text{SOR}^4$, $-\text{SO}_2\text{R}^4$, $-\text{SO}_2\text{NR}^1\text{R}^2$, $-\text{COR}^1$, $-\text{CO}_2\text{R}^1$, $-\text{CONR}^1\text{R}^2$, $-\text{C}(=\text{NR}^1)\text{R}^2$, or $-\text{C}(=\text{NOR}^1)\text{R}^2$ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the $-\text{C}_{1-6}\text{alkyl}$ substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, $-\text{CN}$, $-\text{C}_{1-6}\text{alkyl}$, $-\text{O}(\text{C}_{0-6}\text{alkyl})$, $-\text{O}(\text{C}_{3-7}\text{cycloalkyl})$, $-\text{O}(\text{aryl})$, $-\text{O}(\text{heteroaryl})$, $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{C}_{0-6}\text{alkyl})$, $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{C}_{3-7}\text{cycloalkyl})$, or $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{aryl})$ groups;

R^1 , R^2 , and R^3 each independently is $-\text{C}_{0-6}\text{alkyl}$, $-\text{C}_{3-7}\text{cycloalkyl}$, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, $-\text{CN}$, $-\text{C}_{1-6}\text{alkyl}$, $-\text{O}(\text{C}_{0-6}\text{alkyl})$, $-\text{O}(\text{C}_{3-7}\text{cycloalkyl})$, $-\text{O}(\text{aryl})$, $-\text{O}(\text{heteroaryl})$, $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{C}_{0-6}\text{alkyl})$, $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{C}_{3-7}\text{cycloalkyl})$, $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{aryl})$ substituents;

R^4 is $-\text{C}_{1-6}\text{alkyl}$, $-\text{C}_{3-7}\text{cycloalkyl}$, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, $-\text{CN}$, $-\text{C}_{1-6}\text{alkyl}$, $-\text{O}(\text{C}_{0-6}\text{alkyl})$, $-\text{O}(\text{C}_{3-7}\text{cycloalkyl})$, $-\text{O}(\text{aryl})$, $-\text{O}(\text{heteroaryl})$, $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{C}_{0-6}\text{alkyl})$, $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{C}_{3-7}\text{cycloalkyl})$, $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{aryl})$ substituents;

A is $-C_{0-4}alkyl$, $-C_{0-2}alkyl-SO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-SO_2-C_{0-2}alkyl-$, $-C_{0-2}alkyl-CO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-NR^9CO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-NR^9SO_2-C_{0-2}alkyl-$ or $-heteroC_{0-4}alkyl$;

Y is optionally substituted with 1-7 independent halogen, $-CN$, NO_2 ,
 5 $-C_{1-6}alkyl$, $-C_{1-6}alkenyl$, $-C_{1-6}alkynyl$, $-OR^5$, $-NR^5R^6$, $-C(=NR^5)NR^6R^7$,
 $-N(=NR^5)NR^6R^7$, $-NR^5COR^6$, $-NR^5CO_2R^6$, $-NR^5SO_2R^8$, $-NR^5CONR^6R^7$, $-SR^8$,
 $-SOR^8$, $-SO_2R^8$, $-SO_2NR^5R^6$, $-COR^5$, $-CO_2R^5$, $-CONR^5R^6$, $-C(=NR^5)R^6$, or $-C(=NOR^5)R^6$ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the $-C_{1-6}alkyl$ substituent,
 10 cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, $-CN$, $-C_{1-6}alkyl$, $-O(C_{0-6}alkyl)$, $-O(C_{3-7}cycloalkyl)$, $-O(aryl)$, $-O(heteroaryl)$, $-N(C_{0-6}alkyl)(C_{0-6}alkyl)$, $-N(C_{0-6}alkyl)(C_{3-7}cycloalkyl)$, or $-N(C_{0-6}alkyl)(aryl)$ groups;

R^5 , R^6 , and R^7 each independently is $-C_{0-6}alkyl$, $-C_{3-7}cycloalkyl$,
 15 heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, $-CN$, $-C_{1-6}alkyl$, $-O(C_{0-6}alkyl)$, $-O(C_{3-7}cycloalkyl)$, $-O(aryl)$, $-O(heteroaryl)$, $-N(C_{0-6}alkyl)(C_{0-6}alkyl)$, $-N(C_{0-6}alkyl)(C_{3-7}cycloalkyl)$, $-N(C_{0-6}alkyl)(aryl)$ substituents;

R^8 is $-C_{1-6}alkyl$, $-C_{3-7}cycloalkyl$, heteroaryl, or aryl; optionally
 20 substituted with 1-5 independent halogen, $-CN$, $-C_{1-6}alkyl$, $-O(C_{0-6}alkyl)$, $-O(C_{3-7}cycloalkyl)$, $-O(aryl)$, $-O(heteroaryl)$, $-N(C_{0-6}alkyl)(C_{0-6}alkyl)$, $-N(C_{0-6}alkyl)(C_{3-7}cycloalkyl)$, $-N(C_{0-6}alkyl)(aryl)$ substituents;

B is $-C_{0-4}alkyl$, $-C_{0-2}alkyl-SO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-SO_2-C_{0-2}alkyl-$, $-C_{0-2}alkyl-CO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-NR^{10}CO-C_{0-2}alkyl-$, $-C_{0-2}alkyl-NR^{10}SO_2-C_{0-2}alkyl-$ or $-heteroC_{0-4}alkyl$;

R^9 and R^{10} each independently is $-C_{0-6}alkyl$, $-C_{3-7}cycloalkyl$,
 heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent halogen, $-CN$, $-C_{1-6}alkyl$, $-O(C_{0-6}alkyl)$, $-O(C_{3-7}cycloalkyl)$, $-O(aryl)$, $-O(heteroaryl)$, $-N(C_{0-6}alkyl)(C_{0-6}alkyl)$, $-N(C_{0-6}alkyl)(C_{3-7}cycloalkyl)$, $-N(C_{0-6}alkyl)(aryl)$ substituents; and
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any N may be an N-oxide.

2. The compound according to Claim 1, or a pharmaceutically acceptable salt thereof, wherein

X is 2-pyridyl optionally substituted with 1-4 independent halogen, -CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁-6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups.

3. The compound according to Claim 2, or a pharmaceutically acceptable salt thereof, wherein

Y is phenyl optionally substituted with 1-5 independent halogen, -CN, -NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁-6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups..

4. The compound according to Claim 1, or a pharmaceutically acceptable salt thereof, wherein

Y is 2-pyridyl optionally substituted with 1-4 independent halogen, -CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁-6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further

substituted with 1-5 independent halogen, $-\text{CN}$, $-\text{C}_{1-6}\text{alkyl}$, $-\text{O}(\text{C}_{0-6}\text{alkyl})$, $-\text{O}(\text{C}_{3-7}\text{cycloalkyl})$, $-\text{O}(\text{aryl})$, $-\text{O}(\text{heteroaryl})$, $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{C}_{0-6}\text{alkyl})$, $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{C}_{3-7}\text{cycloalkyl})$, or $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{aryl})$ groups.

5 5. The compound according to Claim 1, or a pharmaceutically acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent halogen, $-\text{CN}$, NO_2 , $-\text{C}_{1-6}\text{alkyl}$, $-\text{OR}^1$, $-\text{NR}^1\text{R}^2$, $-\text{C}(=\text{NR}^1)\text{NR}^2\text{R}^3$, $-\text{N}(=\text{NR}^1)\text{NR}^2\text{R}^3$, $-\text{NR}^1\text{COR}^2$, $-\text{NR}^1\text{CO}_2\text{R}^2$, $-\text{NR}^1\text{SO}_2\text{R}^4$, $-\text{NR}^1\text{CONR}^2\text{R}^3$, $-\text{SR}^4$, $-\text{SOR}^4$, $-\text{SO}_2\text{R}^4$, $-\text{SO}_2\text{NR}^1\text{R}^2$, $-\text{COR}^1$, $-\text{CO}_2\text{R}^1$, $-\text{CONR}^1\text{R}^2$, $-\text{C}(=\text{NR}^1)\text{R}^2$, or $-\text{C}(=\text{NOR}^1)\text{R}^2$ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the $-\text{C}_{1-6}\text{alkyl}$ substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, $-\text{CN}$, $-\text{C}_{1-6}\text{alkyl}$, $-\text{O}(\text{C}_{0-6}\text{alkyl})$, $-\text{O}(\text{C}_{3-7}\text{cycloalkyl})$, $-\text{O}(\text{aryl})$, $-\text{O}(\text{heteroaryl})$, $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{C}_{0-6}\text{alkyl})$, $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{C}_{3-7}\text{cycloalkyl})$, or $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{aryl})$ groups.

6. The compound according to Claim 5, or a pharmaceutically acceptable salt thereof, wherein

20 Y is 2-pyridyl optionally substituted with 1-4 independent halogen, $-\text{CN}$, NO_2 , $-\text{C}_{1-6}\text{alkyl}$, $-\text{C}_{1-6}\text{alkenyl}$, $-\text{C}_{1-6}\text{alkynyl}$, $-\text{OR}^5$, $-\text{NR}^5\text{R}^6$, $-\text{C}(=\text{NR}^5)\text{NR}^6\text{R}^7$, $-\text{N}(=\text{NR}^5)\text{NR}^6\text{R}^7$, $-\text{NR}^5\text{COR}^6$, $-\text{NR}^5\text{CO}_2\text{R}^6$, $-\text{NR}^5\text{SO}_2\text{R}^8$, $-\text{NR}^5\text{CONR}^6\text{R}^7$, $-\text{SR}^8$, $-\text{SOR}^8$, $-\text{SO}_2\text{R}^8$, $-\text{SO}_2\text{NR}^5\text{R}^6$, $-\text{COR}^5$, $-\text{CO}_2\text{R}^5$, $-\text{CONR}^5\text{R}^6$, $-\text{C}(=\text{NR}^5)\text{R}^6$, or $-\text{C}(=\text{NOR}^5)\text{R}^6$ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the $-\text{C}_{1-6}\text{alkyl}$ substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, $-\text{CN}$, $-\text{C}_{1-6}\text{alkyl}$, $-\text{O}(\text{C}_{0-6}\text{alkyl})$, $-\text{O}(\text{C}_{3-7}\text{cycloalkyl})$, $-\text{O}(\text{aryl})$, $-\text{O}(\text{heteroaryl})$, $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{C}_{0-6}\text{alkyl})$, $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{C}_{3-7}\text{cycloalkyl})$, or $-\text{N}(\text{C}_{0-6}\text{alkyl})(\text{aryl})$ groups..

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7. The compound according to Claim 1, or a pharmaceutically acceptable salt thereof, wherein

Y is 1,3-thiazol-2-yl optionally substituted with 1-2 independent halogen, $-\text{CN}$, NO_2 , $-\text{C}_{1-6}\text{alkyl}$, $-\text{C}_{1-6}\text{alkenyl}$, $-\text{C}_{1-6}\text{alkynyl}$, $-\text{OR}^5$, $-\text{NR}^5\text{R}^6$,

-C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁-6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups.

8. The compound according to Claim 7, or a pharmaceutically acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent halogen, -CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁-6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups.

9. The compound according to Claim 1, or a pharmaceutically acceptable salt thereof, wherein

X is 3-pyridyl optionally substituted with 1-4 independent halogen, -CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁-6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups.

10. The compound according to Claim 1, or a pharmaceutically acceptable salt thereof, wherein

Y is imidazolyl optionally substituted with 1-2 independent halogen, –
 5 CN, NO₂, –C₁-6alkyl, –C₁-6alkenyl, –C₁-6alkynyl, –OR⁵, –NR⁵R⁶,
 –C(=NR⁵)NR⁶R⁷, –N(=NR⁵)NR⁶R⁷, –NR⁵COR⁶, –NR⁵CO₂R⁶, –NR⁵SO₂R⁸, –
 NR⁵CONR⁶R⁷, –SR⁸, –SOR⁸, –SO₂R⁸, –SO₂NR⁵R⁶, –COR⁵, –CO₂R⁵, –CONR⁵R⁶,
 –C(=NR⁵)R⁶, or –C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are
 combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the –C₁-
 10 6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further
 substituted with 1-5 independent halogen, –CN, –C₁-6alkyl, –O(C₀-6alkyl), –O(C₃-
 7cycloalkyl), –O(aryl), –O(heteroaryl), –N(C₀-6alkyl)(C₀-6alkyl), –N(C₀-6alkyl)(C₃-
 7cycloalkyl), or –N(C₀-6alkyl)(aryl) groups.

15 11. The compound according to Claim 10, or a pharmaceutically acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent halogen, –CN,
 NO₂, –C₁-6alkyl, –C₁-6alkenyl, –C₁-6alkynyl, –OR¹, –NR¹R², –C(=NR¹)NR²R³, –
 N(=NR¹)NR²R³, –NR¹COR², –NR¹CO₂R², –NR¹SO₂R⁴, –NR¹CONR²R³, –SR⁴,
 20 –SOR⁴, –SO₂R⁴, –SO₂NR¹R², –COR¹, –CO₂R¹, –CONR¹R², –C(=NR¹)R², or –
 C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form
 a cycloalkyl or heterocycloalkyl ring fused to X; wherein the –C₁-6alkyl substituent,
 cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5
 independent halogen, –CN, –C₁-6alkyl, –O(C₀-6alkyl), –O(C₃-7cycloalkyl), –
 25 O(aryl), –O(heteroaryl), –N(C₀-6alkyl)(C₀-6alkyl), –N(C₀-6alkyl)(C₃-7cycloalkyl), or
 –N(C₀-6alkyl)(aryl) groups.

12. The compound according to Claim 1, or a pharmaceutically acceptable salt thereof, wherein

30 Y is 1,3-oxazolyl optionally substituted with 1-2 independent halogen,
 –CN, NO₂, –C₁-6alkyl, –C₁-6alkenyl, –C₁-6alkynyl, –OR⁵, –NR⁵R⁶,
 –C(=NR⁵)NR⁶R⁷, –N(=NR⁵)NR⁶R⁷, –NR⁵COR⁶, –NR⁵CO₂R⁶, –NR⁵SO₂R⁸, –
 NR⁵CONR⁶R⁷, –SR⁸, –SOR⁸, –SO₂R⁸, –SO₂NR⁵R⁶, –COR⁵, –CO₂R⁵, –CONR⁵R⁶,
 –C(=NR⁵)R⁶, or –C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are
 35 combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the –C₁-

6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups.

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13. The compound according to Claim 12, or a pharmaceutically acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent halogen, -CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁-6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups.

14. The compound according to Claim 1, or a pharmaceutically acceptable salt thereof, wherein

Y is isoxazolyl optionally substituted with 1-2 independent halogen, -CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁-6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups.

15. The compound according to Claim 14, or a pharmaceutically acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent halogen, -CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁-6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups.

16. The compound according to Claim 1, or a pharmaceutically acceptable salt thereof, wherein

Y is pyrazolyl optionally substituted with 1-3 independent halogen, -CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁-6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups.

17. The compound according to Claim 16, or a pharmaceutically acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent halogen, -CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁-6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -

O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups.

18. The compound according to Claim 1, or a pharmaceutically acceptable salt thereof, wherein

Y is triazolyl optionally substituted with 1-2 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶, -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups.

19. The compound according to Claim 18, or a pharmaceutically acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups.

20. The compound according to Claim 1, or a pharmaceutically acceptable salt thereof, wherein

Y is pyrimidinyl optionally substituted with 1-3 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR⁵, -NR⁵R⁶, -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -

NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶,
 -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are
 combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁-
 6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further
 5 substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-
 7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-
 7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups.

21. The compound according to Claim 20, or a pharmaceutically
 10 acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent halogen, -CN,
 NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -
 N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴,
 -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -
 15 C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form
 a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁-6alkyl substituent,
 cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5
 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -
 O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or
 20 -N(C₀-6alkyl)(aryl) groups.

22. The compound according to Claim 1, or a pharmaceutically
 acceptable salt thereof, wherein

Y is pyrazinyl optionally substituted with 1-3 independent halogen, -
 25 CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR⁵, -NR⁵R⁶,
 -C(=NR⁵)NR⁶R⁷, -N(=NR⁵)NR⁶R⁷, -NR⁵COR⁶, -NR⁵CO₂R⁶, -NR⁵SO₂R⁸, -
 NR⁵CONR⁶R⁷, -SR⁸, -SOR⁸, -SO₂R⁸, -SO₂NR⁵R⁶, -COR⁵, -CO₂R⁵, -CONR⁵R⁶,
 -C(=NR⁵)R⁶, or -C(=NOR⁵)R⁶ substituents, wherein optionally two substituents are
 combined to form a cycloalkyl or heterocycloalkyl ring fused to Y; wherein the -C₁-
 30 6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further
 substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-
 7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-
 7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups.

23. The compound according to Claim 22, or a pharmaceutically acceptable salt thereof, wherein

X is phenyl optionally substituted with 1-5 independent halogen, -CN, NO₂, -C₁-6alkyl, -C₁-6alkenyl, -C₁-6alkynyl, -OR¹, -NR¹R², -C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁-6alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further substituted with 1-5 independent halogen, -CN, -C₁-6alkyl, -O(C₀-6alkyl), -O(C₃-7cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀-6alkyl)(C₀-6alkyl), -N(C₀-6alkyl)(C₃-7cycloalkyl), or -N(C₀-6alkyl)(aryl) groups.

24. The compound according to Claim 1, consisting of

2-[2-(3-chlorophenyl)-2*H*-tetrazol-5-yl]pyridine;

3-(5-pyridin-2-yl-2*H*-tetrazol-2-yl)benzonitrile;

2-(2-pyridin-3-yl-2*H*-tetrazol-5-yl)pyridine;

2-(3-chlorophenyl)-5-(2-methyl-1,3-thiazol-4-yl)-2*H*-tetrazole;

3-[5-(2-methyl-1,3-thiazol-4-yl)-2*H*-tetrazol-2-yl]benzonitrile;

2-[5-(3-bromophenyl)-2*H*-tetrazol-2-yl]pyridine;

2-[5-(3-chlorophenyl)-2*H*-tetrazol-2-yl]pyridine;

3-(2-pyridin-2-yl-2*H*-tetrazol-5-yl)benzonitrile;

2-[2-(3,5-difluorophenyl)-2*H*-tetrazol-5-yl]pyridine;

2-[2-(3-methoxyphenyl)-2*H*-tetrazol-5-yl]pyridine;

2-[2-(3-trifluoromethylphenyl)-2*H*-tetrazol-5-yl]pyridine;

2-[2-(3-iodophenyl)-2*H*-tetrazol-5-yl]pyridine;

2-[2-(3-bromophenyl)-2*H*-tetrazol-5-yl]pyridine;

2-[2-(3-Methylmercaptophenyl)-2*H*-tetrazol-5-yl]pyridine;

2-[2-(4-fluorophenyl)-2*H*-tetrazol-5-yl]pyridine;

2-[2-(3-fluorophenyl)-2*H*-tetrazol-5-yl]pyridine;

2-[2-(2-methoxyphenyl)-2*H*-tetrazol-5-yl]pyridine;

2-[2-(3-ethylphenyl)-2*H*-tetrazol-5-yl]pyridine;

2-[2-(3-methylphenyl)-2*H*-tetrazol-5-yl]pyridine;

2-[2-(2-chloro-3-pyridyl)-2*H*-tetrazol-5-yl]pyridine;

2-[2-(3,5-dichlorophenyl)-2*H*-tetrazol-5-yl]pyridine;

- 2-[2-(2-chlorophenyl)-2*H*-tetrazol-5-yl]pyridine;
 2-[2-(4-methoxyphenyl)-2*H*-tetrazol-5-yl]pyridine;
 2-[2-(4-pyridyl)-2*H*-tetrazol-5-yl]pyridine;
 2-[2-(3,5-dimethylphenyl)-2*H*-tetrazol-5-yl]pyridine;
 5 3-[5-(6-methylpyridin-2-yl)-2*H*-tetrazol-2-yl]benzonitrile;
 3-[5-(4-methylpyridin-2-yl)-2*H*-tetrazol-2-yl]benzonitrile;
 3-[5-(4-methylpyridin-2-yl)-2*H*-tetrazol-2-yl]benzonitrile;
 2-[2-(3-isopropylphenyl)-2*H*-tetrazol-5-yl]pyridine;
 2-{2-[3-(trifluoromethoxy)phenyl]-2*H*-tetrazol-5-yl}pyridine;
 10 2-[2-(3-ethoxyphenyl)-2*H*-tetrazol-5-yl]pyridine;
 2-{2-[3-methoxy-5-(trifluoromethyl)phenyl]-2*H*-tetrazol-5-
 yl}pyridine;
 3-[5-(1,3-thiazol-2-yl)-2*H*-tetrazol-2-yl]benzonitrile;
 3-[5-(5-methylpyridin-2-yl)-2*H*-tetrazol-2-yl]benzonitrile;
 15 3-[5-(3-methylpyridin-2-yl)-2*H*-tetrazol-2-yl]benzonitrile;
 3-[5-(1-methyl-1*H*-imidazol-2-yl)-2*H*-tetrazol-2-yl]benzonitrile;
 2-[2-(3,4-dimethylphenyl)-2*H*-tetrazol-5-yl]pyridine;
 2-[2-(3,4,5-trichlorophenyl)-2*H*-tetrazol-5-yl]pyridine;
 3-[5-(1,3-thiazol-4-yl)-2*H*-tetrazol-2-yl]benzonitrile;
 20 2-[2-(2,5-dichlorophenyl)-2*H*-tetrazol-5-yl]pyridine;
 2-[2-(3,4-difluorophenyl)-2*H*-tetrazol-5-yl]pyridine;
 2-[2-(3-chloro-4-methylphenyl)-2*H*-tetrazol-5-yl]pyridine;
 2-[2-(2,3-dichlorophenyl)-2*H*-tetrazol-5-yl]pyridine;
 2-{2-[3-methyl-5-(trifluoromethyl)phenyl]-2*H*-tetrazol-5-yl}pyridine;
 25 2-[2-(3-bromo-4-methylphenyl)-2*H*-tetrazol-5-yl]pyridine;
 2-[2-(3-chloro-4-iodophenyl)-2*H*-tetrazol-5-yl]pyridine;
 2-{2-[3-fluoro-5-(trifluoromethyl)phenyl]-2*H*-tetrazol-5-yl}pyridine;
 3-(5-pyrazin-2-yl-2*H*-tetrazol-2-yl)benzonitrile;
 2-[2-(2,3-dihydro-1*H*-inden-5-yl)-2*H*-tetrazol-5-yl]pyridine;
 30 2-[2-(1,3-dihydro-2-benzofuran-5-yl)-2*H*-tetrazol-5-yl]pyridine;
 2-[2-(4-methoxy-2-naphthyl)-2*H*-tetrazol-5-yl]pyridine;
 2-[2-(1-naphthyl)-2*H*-tetrazol-5-yl]pyridine;
 2-[2-(3,5-dimethoxyphenyl)-2*H*-tetrazol-5-yl]pyridine;
 2-[2-(4-phenoxyphenyl)-2*H*-tetrazol-5-yl]pyridine;
 35 3-[5-(3-fluoro-pyridin-2-yl)-tetrazol-2-yl]-benzonitrile

- 3-(5-isoxazol-3-yl-tetrazol-2-yl)-benzonitrile
- 3-[5-(1-methyl-1H-pyrazol-3-yl)-tetrazol-2-yl]-benzonitrile
- 3-(5-oxazol-4-yl-tetrazol-2-yl)-benzonitrile
- 3-[5-(2H-pyrazol-3-yl)-tetrazol-2-yl]-benzonitrile
- 5 3-[5-(1H-imidazol-2-yl)-tetrazol-2-yl]-benzonitrile
- 3-[5-(5-hydroxy-pyridin-2-yl)-tetrazol-2-yl]-benzonitrile
- 3-[5-(5-bromo-pyridin-2-yl)-tetrazol-2-yl]-5-fluoro-benzonitrile
- 3-fluoro-5-[5-(3-fluoro-pyridin-2-yl)-tetrazol-2-yl]-benzonitrile
- 3-fluoro-5-(5-oxazol-2-yl-tetrazol-2-yl)-benzonitrile
- 10 3-{3-fluoro-5-[5-(1H-imidazol-2-yl)-tetrazol-2-yl]-phenoxy}-pyridine
- 3-fluoro-5-[5-(4-methyl-1H-imidazol-2-yl)-tetrazol-2-yl]-benzonitrile
- 3-{3-fluoro-5-[5-(1H-[1,2,3]triazol-4-yl)-tetrazol-2-yl]-phenoxy}-pyridine
- 2-{2-[3-fluoro-5-(pyridin-3-yloxy)-phenyl]-2H-tetrazol-5-yl}-pyrimidine
- 15 3-{3-fluoro-5-[5-(1H-pyrazol-3-yl)-tetrazol-2-yl]-phenoxy}-pyridine
- 3-{3-fluoro-5-[5-(1-methyl-1H-pyrazol-3-yl)-tetrazol-2-yl]-phenoxy}-pyridine
- 3-fluoro-5-[5-(1-methyl-1H-pyrazol-3-yl)-tetrazol-2-yl]-benzonitrile
- 20 *N*-[4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]pyridin-3-amine
- 2-{2-[4-(pyridin-3-ylmethyl)phenyl]-2*H*-tetraazol-5-yl}pyridine
- N*-[3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]pyridin-3-amine
- N*-[3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]pyridin-2-amine
- 2-{2-[3-(pyridin-2-yloxy)phenyl]-2*H*-tetraazol-5-yl}pyridine
- 25 2-[2-(3-ethynylphenyl)-2*H*-tetraazol-5-yl]pyridine
- 2-{2-[4-(pyridin-3-yloxy)phenyl]-2*H*-tetraazol-5-yl}pyridine
- 2-{2-[3-(pyridin-3-yloxy)phenyl]-2*H*-tetraazol-5-yl}pyridine
- 2-[2-(3-nitrophenyl)-2*H*-tetraazol-5-yl]pyridine
- 2-methyl-3-[3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenoxy]pyridine
- 30 2-methyl-*N*-[3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]pyridin-3-amine
- N*-methyl-*N*-[3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]pyridin-3-amine
- N*-[3-fluoro-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]-*N*-methylpyridin-3-amine
- 35

- 2-{2-[3-fluoro-5-(pyridin-3-yloxy)phenyl]-2*H*-tetraazol-5-yl}pyridine
N-[3-fluoro-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]pyridin-3-amine
5 *N*-{3-[5-(5-bromopyridin-2-yl)-2*H*-tetraazol-2-yl]-5-fluorophenyl}pyridin-3-amine
2-{2-[3-fluoro-5-(pyridin-3-ylmethyl)phenyl]-2*H*-tetraazol-5-yl}pyridine
5-[3-fluoro-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)benzyl]-2-methylpyridine
10 3-{3-fluoro-5-[5-(1,3-thiazol-2-yl)-2*H*-tetraazol-2-yl]phenoxy}pyridine
2-{2-[3-fluoro-5-(pyridin-3-ylthio)phenyl]-2*H*-tetraazol-5-yl}pyridine
2-(2-{3-fluoro-5-[(pyridin-3-yloxy)methyl]phenyl}-2*H*-tetraazol-5-yl)pyridine
3-fluoro-2-{2-[3-fluoro-5-(pyridin-3-yloxy)phenyl]-2*H*-tetraazol-5-yl}pyridine
15 2-{2-[3-fluoro-5-(pyridin-3-yloxy)phenyl]-2*H*-tetraazol-5-yl}-6-methylpyridine
2-{2-[3-methoxy-4-(3*H*-1,2λ⁵,3,4-tetraazol-2-ylmethyl)phenyl]-2*H*-tetraazol-5-yl}pyridine
20 3-bromo-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)pyridine
2-{2-[4-(allyloxy)-3-methoxyphenyl]-2*H*-tetraazol-5-yl}pyridine
2-methoxy-4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenol
2-methoxy-4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl ethylcarbamate
2-[2-(4-bromo-3-methoxyphenyl)-2*H*-tetraazol-5-yl]pyridine
25 2-{2-[3-methoxy-4-(pyridin-2-yloxy)phenyl]-2*H*-tetraazol-5-yl}pyridine
5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)nicotinonitrile
[3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenoxy]acetonitrile
N-pyridin-3-yl-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)pyridin-3-amine
30 3-(pyridin-3-yloxy)-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)pyridine
3-bromo-5-[5-(1*H*-imidazol-2-yl)-2*H*-tetraazol-2-yl]pyridine
3-bromo-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)pyridine
2-(pyridin-3-yloxy)-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)pyridine
2-[(5-chloropyridin-3-yl)oxy]-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)pyridine
35

- 2-methyl-5-{[5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)pyridin-2-yl]oxy}pyridine
- 2-methyl-3-{[5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)pyridin-2-yl]oxy}pyridine
- 5 2-[(4-methylpyridin-3-yl)oxy]-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)pyridine
- 2-(pyridin-4-yloxy)-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)pyridine
- 2-bromo-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)benzonitrile
- 3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)aniline
- 10 3-fluoro-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)benzonitrile
- 2-{2-[3-bromo-5-(trifluoromethyl)phenyl]-2*H*-tetraazol-5-yl}pyridine
- 3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)-5-(trifluoromethyl)benzonitrile
- 3-nitro-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)benzonitrile
- 3-amino-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)benzonitrile
- 15 3-chloro-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)benzonitrile
- 2-[2-(3-cyano-5-fluorophenyl)-2*H*-tetraazol-5-yl]-1-hydroxypyridinium
- 5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)isophthalonitrile
- 2-{5-[3-chloro-5-(pyridin-3-yloxy)phenyl]-2*H*-tetraazol-2-yl}pyridine
- 2-[5-(3-bromo-5-chlorophenyl)-2*H*-tetraazol-2-yl]pyridine
- 20 3-chloro-5-(2-pyridin-2-yl-2*H*-tetraazol-5-yl)benzonitrile
- 3-methyl-5-(2-pyridin-2-yl-2*H*-tetraazol-5-yl)benzonitrile
- [3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]methanol
- [3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]acetonitrile
- N*-methyl-3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)aniline
- 25 2-{2-[3-(1*H*-imidazol-1-ylmethyl)phenyl]-2*H*-tetraazol-5-yl}pyridine
- 2-methoxy-*N*-[3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]aniline
- 2-[2-(3-fluoro-5-iodophenyl)-2*H*-tetraazol-5-yl]pyridine
- 3-fluoro-5-[5-(1*H*-imidazol-2-yl)-2*H*-tetraazol-2-yl]benzonitrile
- 3-[5-(4,5-dibromo-1*H*-imidazol-2-yl)-2*H*-tetraazol-2-yl]-5-
- 30 fluorobenzonitrile
- 2-{2-[3-fluoro-5-(pyridin-3-ylmethoxy)phenyl]-2*H*-tetraazol-5-yl}pyridine
- 2-{[3-fluoro-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenoxy]methyl}benzonitrile

R⁸ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

5 B is -C₀₋₄alkyl, -C₀₋₂alkyl-SO-C₀₋₂alkyl-, -C₀₋₂alkyl-SO₂-C₀₋₂alkyl-, -C₀₋₂alkyl-CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹⁰CO-C₀₋₂alkyl-, -C₀₋₂alkyl-NR¹SO₂-C₀₋₂alkyl- or -heteroC₀₋₄alkyl;

R⁹ and R¹⁰ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent
 10 halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents; and

any N may be an N-oxide.

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In a fourth aspect, the compounds of this invention are represented by Formula (I) or a pharmaceutically acceptable salt thereof, wherein

X is aryl or heteroaryl optionally substituted with 1-7 independent halogen, -CN, NO₂, -C₁₋₆alkyl, -C₁₋₆alkenyl, -C₁₋₆alkynyl, -OR¹, -NR¹R², -
 20 C(=NR¹)NR²R³, -N(=NR¹)NR²R³, -NR¹COR², -NR¹CO₂R², -NR¹SO₂R⁴, -NR¹CONR²R³, -SR⁴, -SOR⁴, -SO₂R⁴, -SO₂NR¹R², -COR¹, -CO₂R¹, -CONR¹R², -C(=NR¹)R², or -C(=NOR¹)R² substituents, wherein optionally two substituents are combined to form a cycloalkyl or heterocycloalkyl ring fused to X; wherein the -C₁₋₆alkyl substituent, cycloalkyl ring, or heterocycloalkyl ring each optionally is further
 25 substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), or -N(C₀₋₆alkyl)(aryl) groups;

R¹, R², and R³ each independently is -C₀₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; any of which is optionally substituted with 1-5 independent
 30 halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋₇cycloalkyl), -O(aryl), -O(heteroaryl), -N(C₀₋₆alkyl)(C₀₋₆alkyl), -N(C₀₋₆alkyl)(C₃₋₇cycloalkyl), -N(C₀₋₆alkyl)(aryl) substituents;

R⁴ is -C₁₋₆alkyl, -C₃₋₇cycloalkyl, heteroaryl, or aryl; optionally substituted with 1-5 independent halogen, -CN, -C₁₋₆alkyl, -O(C₀₋₆alkyl), -O(C₃₋

- N*-{3-fluoro-5-[5-(1*H*-imidazol-2-yl)-2*H*-tetraazol-2-yl]phenyl}-*N*-methylpyridin-3-amine
 3-chloro-5-[5-(1*H*-imidazol-2-yl)-2*H*-tetraazol-2-yl]benzonitrile
 3-{3-fluoro-5-[5-(1-methyl-1*H*-imidazol-2-yl)-2*H*-tetraazol-2-yl]phenoxy}pyridine
 4-chloro-2-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)benzonitrile
 2-[2-(2,4-dichlorophenyl)-2*H*-tetraazol-5-yl]pyridine
 2-[2-(5-chloro-2-methoxyphenyl)-2*H*-tetraazol-5-yl]pyridine
 3-[5-(2-methyl-2*H*-pyrazol-3-yl)-tetrazol-2-yl]-benzonitrile
 3-[5-(1*H*-imidazol-4-yl)-tetrazol-2-yl]-benzonitrile
 3-[5-(5-methoxy-pyridin-2-yl)-tetrazol-2-yl]-benzonitrile
 3-[5-(1*H*-benzoimidazol-2-yl)-tetrazol-2-yl]-benzonitrile
 4-bromo-3-fluoro-5-[5-(3-fluoro-pyridin-2-yl)-tetrazol-2-yl]-benzonitrile
 3-{3-[5-(4,5-dibromo-1*H*-imidazol-2-yl)-tetrazol-2-yl]-5-fluorophenoxy}-pyridine
 7-{2-[3-fluoro-5-(pyridin-3-yloxy)phenyl]-2*H*-tetrazol-5-yl}-pyrazolo[1,5-*a*]pyridine
 2-[2-(5-methylisoxazol-3-yl)-2*H*-tetraazol-5-yl]pyridine
N-phenyl-*N*-[4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]amine
N-[4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]pyridin-2-amine
N-phenyl-*N*-[3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]amine
 2-{2-[4-(pyridin-2-yloxy)phenyl]-2*H*-tetraazol-5-yl}pyridine
 2-{2-[3-(3-methoxyphenoxy)phenyl]-2*H*-tetraazol-5-yl}pyridine
 2-bromo-6-{2-[3-fluoro-5-(pyridin-3-yloxy)phenyl]-2*H*-tetraazol-5-yl}pyridine
 2-{2-[3-fluoro-5-(pyridin-3-yloxy)phenyl]-2*H*-tetraazol-5-yl}-6-methoxypyridine
 2-{2-[3-methoxy-4-(3*H*-1λ⁵,2,3,4-tetraazol-1-ylmethyl)phenyl]-2*H*-tetraazol-5-yl}pyridine
 2-(2-pyridin-3-yl-2*H*-tetraazol-5-yl)pyridine
 2-chloro-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)pyridine
 3-[5-(1*H*-imidazol-2-yl)-2*H*-tetraazol-2-yl]pyridine
 2-(2-pyridin-2-yl-2*H*-tetraazol-5-yl)pyridine
N,N-dimethyl-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)pyridin-2-amine

- N*-pyridin-2-yl-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)pyridin-2-amine
N-pyridin-3-yl-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)pyridin-2-amine
 3-chloro-*N*-[5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)pyridin-2-yl]-5-(trifluoromethyl)pyridin-2-amine
 5 3-(5-quinolin-2-yl-2*H*-tetraazol-2-yl)benzonitrile
 3-(5-isoquinolin-3-yl-2*H*-tetraazol-2-yl)benzonitrile
 2-[2-(4-bromo-3-fluorophenyl)-2*H*-tetraazol-5-yl]pyridine
 2-{2-[3,5-bis(trifluoromethyl)phenyl]-2*H*-tetraazol-5-yl}pyridine
 2-{2-[4-bromo-3-(trifluoromethyl)phenyl]-2*H*-tetraazol-5-yl}pyridine
 10 imethyl 5-(5-pyridin-2-yl-2*H*-tetrazol-2-yl)isophthalate
 2-[3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenoxy]ethylamine
 2-{2-[3-fluoro-5-(pyridin-2-ylmethoxy)phenyl]-2*H*-tetraazol-5-yl}pyridine
 2-[2-(2,6-dichlorophenyl)-2*H*-tetraazol-5-yl]pyridine
 15 [3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]acetic acid
 2-[2-(3-iodo-4-methylphenyl)-2*H*-tetraazol-5-yl]pyridine
 * 2-[2-(4-chloro-3-methylphenyl)-2*H*-tetraazol-5-yl]pyridine
 2-[2-(1,3-benzodioxol-5-yl)-2*H*-tetraazol-5-yl]pyridine
 2-{2-[2-chloro-5-(trifluoromethyl)phenyl]-2*H*-tetraazol-5-yl}pyridine
 20 2-chloro-4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)benzonitrile
 2-[2-(4-bromo-3-methylphenyl)-2*H*-tetraazol-5-yl]pyridine
 2-[2-(3,4-dichlorophenyl)-2*H*-tetraazol-5-yl]pyridine
 2-[2-(3-chloro-4-fluorophenyl)-2*H*-tetraazol-5-yl]pyridine
 2-[2-(3-fluoro-4-methylphenyl)-2*H*-tetraazol-5-yl]pyridine
 25 1-[3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]ethanol
 1-[3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]ethanone
 [3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]methanol
 2-[2-(3-phenoxyphenyl)-2*H*-tetraazol-5-yl]pyridine
 2-{2-[3-(benzyloxy)phenyl]-2*H*-tetraazol-5-yl}pyridine
 30 2-{2-[3-(methylsulfonyl)phenyl]-2*H*-tetraazol-5-yl}pyridine
N-[3-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]acetamide
 2-[2-(3-isopropylphenyl)-2*H*-tetrazol-5-yl]pyridine
 2-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)benzonitrile
 2-(2-phenyl-2*H*-tetraazol-5-yl)pyridine
 35 [2-methoxy-4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]acetonitrile

- 2-[2-(3-methoxy-4-methylphenyl)-2*H*-tetraazol-5-yl]pyridine
- 2-[2-methoxy-4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)-2-methylpropanenitrile
- 2-{cyano[2-methoxy-4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]methyl}nicotinonitrile
- 2-[2-methoxy-4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]succinonitrile
- 2-[2-methoxy-4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]propane-1,2,3-tricarbonitrile
- 2-[2-methoxy-4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]propanenitrile
- 2-{cyano[2-methoxy-4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)methyl]benzonitrile
- (6-fluoropyridin-2-yl)[2-methoxy-4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]acetonitrile
- [2-methoxy-4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl](pyridin-2-yl)acetonitrile
- 2-(2-pyridin-3-yl-2*H*-tetraazol-5-yl)pyridine
- 2-{2-[4-(methylthio)phenyl]-2*H*-tetraazol-5-yl}pyridine
- 2{2-[4-(trifluoromethoxy)phenyl]-2*H*-tetraazol-5-yl}pyridine
- 4-((5-pyridin-2-yl-2*H*-tetraazol-2-yl)benzonitrile
- 2-[2-(4-chlorophenyl)-2*H*-tetraazol-5-yl]pyridine
- 2-[2-(2-naphthyl)-2*H*-tetraazol-5-yl]pyridine
- 2-[2-(3,5-ditert-butylphenyl)-2*H*-tetraazol-5-yl]pyridine
- 2-fluoro-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)benzonitrile
- 2-{2-[4-(bromomethyl)-3-methoxyphenyl]-2*H*-tetraazol-5-yl}pyridine
- 2-methoxy-4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)benzoic acid
- 2-[2-methoxy-4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]ethanamine
- 1-nitroso-6-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)indoline
- 2-[2-methoxy-4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)phenyl]propan-2-ol
- 2-methoxy-4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)benzamide
- 2-methoxy-4-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)benzonitrile
- 3-bromo-6-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)-1*H*-indole
- 6-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)-1*H*-indole
- 6-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)-1*H*-indole-1-carbonitrile

- 5 6-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)-1*H*-indole-3-carbonitrile
 3-methoxy-5-(5-pyridin-2-yl-2*H*-tetraazol-2-yl)benzonitrile
 3-[5-(1*H*-imidazol-2-yl)-2*H*-tetraazol-2-yl]-5-methylbenzonitrile
 3-{3-fluoro-5-[5-(4-methyl-1*H*-imidazol-2-yl)-2*H*-tetraazol-2-yl]phenoxy}pyridine
 3-chloro-5-[5-(1*H*-imidazol-2-yl)-2*H*-tetraazol-2-yl]benzonitrile
- 10 3-methyl-5-[5-(5-methyl-1*H*-imidazol-2-yl)-2*H*-tetraazol-2-yl]benzonitrile
 3-chloro-5-[5-(5-methyl-1*H*-imidazol-2-yl)-2*H*-tetraazol-2-yl]benzonitrile
 3-fluoro-5-[5-(1,3-thiazol-2-yl)-2*H*-tetraazol-2-yl]benzonitrile
 3-fluoro-5-[5-(1,3-thiazol-4-yl)-2*H*-tetraazol-2-yl]benzonitrile
 3-{3-fluoro-5-[5-(1,3-thiazol-4-yl)-2*H*-tetraazol-2-yl]phenoxy}pyridine
 15 3-fluoro-5-{5-[4-(trifluoromethyl)-1*H*-imidazol-2-yl]-2*H*-tetraazol-2-yl}benzonitrile
 3-[5-(4-methyl-1*H*-imidazol-2-yl)-2*H*-tetraazol-2-yl]-5-(pyridin-3-ylloxy)benzonitrile
 3-methoxy-5-[5-(4-methyl-1*H*-imidazol-2-yl)-2*H*-tetraazol-2-yl]benzonitrile
 20

or a pharmaceutically acceptable salt thereof.

- 25 25. A pharmaceutical composition comprising:
 a therapeutically effective amount of the compound according to claim 1, or a pharmaceutically acceptable salt thereof; and
 a pharmaceutically acceptable carrier.

- 30 26. The pharmaceutical composition according to claim 25, further comprising i) an opiate agonist, ii) an opiate antagonist, iii) a calcium channel antagonist, iv) a 5HT receptor agonist, v) a 5HT receptor antagonist, vi) a sodium channel antagonist, vii) an NMDA receptor agonist, viii) an NMDA receptor antagonist, ix) a COX-2 selective inhibitor, x) an NK1 antagonist, xi) a non-steroidal anti-inflammatory drug, xii) a GABA-A receptor modulator, xiii) a dopamine agonist,
 35 xiv) a dopamine antagonist, xv) a selective serotonin reuptake inhibitor, xvi) a

tricyclic antidepressant drug, xvii) a norepinephrine modulator, xviii) L-DOPA, xix) buspirone, xx) a lithium salt, xxi) valproate, xxii) neurontin, xxiii) olanzapine, xxiv) a nicotinic agonist, xxv) a nicotinic antagonist, xxvi) a muscarinic agonist, xxvii) a muscarinic antagonist, xxviii) a selective serotonin and norepinephrine reuptake inhibitor (SSNRI), xxix) a heroin substituting drug, xxx) disulfiram, or xxxi) acamprosate.

27. The pharmaceutical composition according to claim 26, wherein said heroin substituting drug is methadone, levo-alpha-acetylmethadol, buprenorphine or naltrexone.

28. A method of treatment or prevention of pain comprising the step of administering a therapeutically effective amount, or a prophylactically effective amount, of the compound according to claim 1 or a pharmaceutically acceptable salt thereof.

29. A method of treatment or prevention of a pain disorder wherein said pain disorder is acute pain, persistent pain, chronic pain, inflammatory pain, or neuropathic pain, comprising the step of administering a therapeutically effective amount, or a prophylactically effective amount, of the compound according to claim 1 or a pharmaceutically acceptable salt thereof.

30. A method of treatment or prevention of anxiety, depression, bipolar disorder, psychosis, drug withdrawal, tobacco withdrawal, memory loss, cognitive impairment, dementia, Alzheimer's disease, schizophrenia or panic comprising the step of administering a therapeutically effective amount, or a prophylactically effective amount, of the compound according to claim 1 or a pharmaceutically acceptable salt thereof.

31. A method of treatment or prevention of disorders of extrapyramidal motor function comprising the step of administering a therapeutically effective amount, or a prophylactically effective amount, of the compound according to claim 1 or a pharmaceutically acceptable salt thereof.

32. The method of claim 31 wherein said disorder of extrapyramidal motor function is Parkinson's disease, progressive supramuscular palsy, Huntington's disease, Gilles de la Tourette syndrome, or tardive dyskinesia.

5 33. A method of treatment or prevention of anxiety disorders comprising the step of administering a therapeutically effective amount, or a prophylactically effective amount, of the compound according to claim 1 or a pharmaceutically acceptable salt thereof.

10 34. The method of claim 33 wherein said anxiety disorder is panic attack, agoraphobia or specific phobias, obsessive-compulsive disorders, post-traumatic stress disorder, acute stress disorder, generalized anxiety disorder, eating disorder, substance-induced anxiety disorder, or nonspecified anxiety disorder.

15 35. A method of treatment or prevention of neuropathic pain comprising the step of administering a therapeutically effective amount, or a prophylactically effective amount, of the compound according to claim 1 or a pharmaceutically acceptable salt thereof.

20 36. A method of treatment or prevention of Parkinson's Disease comprising the step of administering a therapeutically effective amount, or a prophylactically effective amount, of the compound according to claim 1 or a pharmaceutically acceptable salt thereof.

25 37. A method of treatment or prevention of depression comprising the step of administering a therapeutically effective amount, or a prophylactically effective amount, of the compound according to claim 1 or a pharmaceutically acceptable salt thereof.

30 38. A method of treatment or prevention of epilepsy comprising the step of administering a therapeutically effective amount, or a prophylactically effective amount, of the compound according to claim 1 or a pharmaceutically acceptable salt thereof.

39. A method of treatment or prevention of inflammatory pain comprising the step of administering a therapeutically effective amount, or a prophylactically effective amount, of the compound according to claim 1 or a pharmaceutically acceptable salt thereof.

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40. A method of treatment or prevention of cognitive dysfunction comprising the step of administering a therapeutically effective amount, or a prophylactically effective amount, of the compound according to claim 1 or a pharmaceutically acceptable salt thereof.

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41. A method of treatment or prevention of drug addiction, drug abuse and drug withdrawal comprising the step of administering a therapeutically effective amount, or a prophylactically effective amount, of the compound according to claim 1 or a pharmaceutically acceptable salt thereof.

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42. A method of treatment or prevention of bipolar disorders comprising the step of administering a therapeutically effective amount, or a prophylactically effective amount, of the compound according to claim 1 or a pharmaceutically acceptable salt thereof.

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43. A method of treatment or prevention of circadian rhythm and sleep disorders comprising the step of administering a therapeutically effective amount, or a prophylactically effective amount, of the compound according to claim 1 or a pharmaceutically acceptable salt thereof.

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44. The method of Claim 43 wherein the circadian rhythm and sleep disorders are shift-work induced sleep disorder or jet-lag.

45. A method of treatment or prevention of obesity comprising the step of administering a therapeutically effective amount, or a prophylactically effective amount, of the compound according to claim 1 or a pharmaceutically acceptable salt thereof.

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